WO 05/012335
PCT/GB04/03150

10566487
30 July 2003

Cuthbertson
10/566487
3/15/7 Hughs Search

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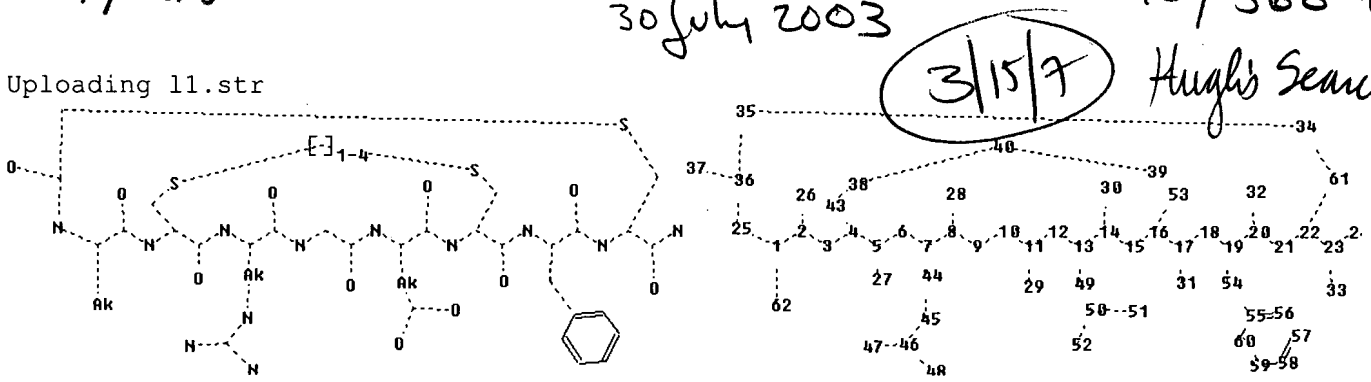

chain nodes :
23 24 26 27 28 29 30 31 32 33 37 44 45 46 47 48 49 50 51 52 54
62
ring nodes :
1 2 3 4 5 6 7 8 9 10 11 12 13 14 15 16 17 18 19 20 21 22 25
34 35 36 38 39 40 43 53 55 56 57 58 59 60 61
chain bonds :
1-62 2-26 5-27 7-44 8-28 11-29 13-49 14-30 17-31 19-54 20-32 22-23 23-
24
23-33 36-37 44-45 45-46 46-47 46-48 49-50 50-51 50-52 54-55
ring bonds :
1-2 1-25 2-3 3-4 4-5 4-43 5-6 6-7 7-8 8-9 9-10 10-11 11-12 12-13 13-
14
14-15 15-16 16-17 16-53 17-18 18-19 19-20 20-21 21-22 22-61 25-36 34-35
34-61 35-36
38-40 38-43 39-40 39-53 55-56 55-60 56-57 57-58 58-59 59-60
exact/norm bonds :
1-2 1-25 1-62 2-3 2-26 3-4 4-5 4-43 5-6 5-27 6-7 7-8 7-44 8-9 8-28
9-10 10-11 11-12 11-29 12-13 13-14 13-49 14-15 14-30 15-16 16-17 16-53
17-18 17-31 18-19
19-20 19-54 20-21 20-32 21-22 22-23 22-61 23-24 23-33 25-36 34-35 34-61
35-36 36-37
38-40 38-43 39-40 39-53 44-45 45-46 46-47 46-48 49-50 50-51 50-52 54-55
normalized bonds :
55-56 55-60 56-57 57-58 58-59 59-60

Match level :
1:Atom 2:Atom 3:Atom 4:Atom 5:Atom 6:Atom 7:Atom 8:Atom 9:Atom 10:Atom
11:Atom 12:Atom 13:Atom 14:Atom 15:Atom 16:Atom 17:Atom 18:Atom 19:Atom
20:Atom 21:Atom
22:Atom 23:CLASS 24:CLASS 25:Atom 26:CLASS 27:CLASS 28:CLASS 29:CLASS
30:CLASS 31:CLASS 32:CLASS
33:CLASS 34:Atom 35:Atom 36:Atom 37:CLASS 38:Atom 39:Atom 40:Atom 43:Atom
44:CLASS
45:CLASS 46:CLASS 47:CLASS 48:CLASS 49:CLASS 50:CLASS 51:CLASS 52:CLASS
53:Atom 54:CLASS
55:Atom 56:Atom 57:Atom 58:Atom 59:Atom 60:Atom 61:Atom 62:CLASS

=> d que l21
L15 94 SEA FILE=HCAPLUS ABB=ON PLU=ON ("CUTHBERTSON A"/AU OR

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                "CUTHBERTSON A C"/AU OR "CUTHBERTSON A F"/AU OR "CUTHBERTSON A
                F J"/AU OR "CUTHBERTSON A G S"/AU OR "CUTHBERTSON A M"/AU OR
                "CUTHBERTSON A S"/AU OR "CUTHBERTSON A Z"/AU OR "CUTHBERTSON
                ALAN"/AU OR "CUTHBERTSON ALAN J S"/AU OR "CUTHBERTSON ALAN
                S"/AU OR "CUTHBERTSON ALLAN S"/AU)
L16          28 SEA FILE=HCAPLUS ABB=ON  PLU=ON  ("SOLBAKKEN M"/AU OR "SOLBAKKE
                N MAGNE"/AU)
L17          19 SEA FILE=HCAPLUS ABB=ON  PLU=ON  L15 AND L16
L18          14 SEA FILE=HCAPLUS ABB=ON  PLU=ON  L17 AND (AY<2004 OR PY<2004
                OR PRY<2004)
L19          32 SEA FILE=HCAPLUS ABB=ON  PLU=ON  (L15 OR L16) AND (IMAGING?)
L20          20 SEA FILE=HCAPLUS ABB=ON  PLU=ON  L19 AND (AY<2004 OR PY<2004
                OR PRY<2004)
L21          24 SEA FILE=HCAPLUS ABB=ON  PLU=ON  (L20 OR L18)


=> d que l30
L22         437 SEA CUTHBERTSON A?/AU
L23          74 SEA SOLBAKKEN M?/AU
L24          49 SEA L22 AND L23
L25          36 SEA L24 AND IMAGING?
L26          25 SEA L25 AND (AY<2004 OR PY<2004 OR PRY<2004)
L27          47 SEA (L22 OR L23) AND (IMAGING AGENT?)
L28          29 SEA L27 AND (AY<2004 OR PY<2004 OR PRY<2004)
L29          40 SEA (L26 OR L28)
L30          29 SEA L29 AND (IMAGING AGENT?)


=> dup rem l21,l30
FILE 'HCAPLUS' ENTERED AT 13:57:33 ON 15 MAR 2007
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PROCESSING COMPLETED FOR L30
L36          25 DUP REM L21 L30 (28 DUPLICATES REMOVED)
                ANSWERS '1-24' FROM FILE HCAPLUS
                ANSWER '25' FROM FILE WPIX

=> d ibib abs hitstr retable l36 tot

L36  ANSWER 1 OF 25  HCAPLUS  COPYRIGHT 2007 ACS on STN DUPLICATE 1
ACCESSION NUMBER:         2005:472005  HCAPLUS  Full-text
DOCUMENT NUMBER:          143:13254
TITLE:                    Contrast agent
INVENTOR(S):              Cuthbertson, Alan; Solbakken, Magne
                          ; Lovhaug, Dagfinn
PATENT ASSIGNEE(S):       Amersham Health AS, Norway
SOURCE:                   PCT Int. Appl., 30 pp.
```

CODEN: PIXXD2
DOCUMENT TYPE: Patent
LANGUAGE: English
FAMILY ACC. NUM. COUNT: 3
PATENT INFORMATION:

| PATENT NO. | KIND | DATE | APPLICATION NO. | DATE |
|---|---|---|---|---|
| WO 2005049095 | A2 | 20050602 | WO 2004-NO358 | 20041123 <-- |
| WO 2005049095 | A3 | 20060112 | | |
| W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BW, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, EG, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NA, NI, NO, NZ, OM, PG, PH, PL, PT, RO, RU, SC, SD, SE, SG, SK, SL, SY, TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, YU, ZA, ZM, ZW | | | | |
| RW: BW, GH, GM, KE, LS, MW, MZ, NA, SD, SL, SZ, TZ, UG, ZM, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM, AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE, IS, IT, LU, MC, NL, PL, PT, RO, SE, SI, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG | | | | |
| EP 1699494 | A2 | 20060913 | EP 2004-808852 | 20041123 <-- |
| R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, FI, RO, CY, TR, BG, CZ, EE, HU, PL, SK, IS | | | | |
| US 2006193768 | A1 | 20060831 | US 2005-559880 | 20051207 <-- |

PRIORITY APPLN. INFO.: NO 2003-5228 A 20031124 <--
GB 2004-16062 A 20040719
WO 2004-NO358 W 20041123

OTHER SOURCE(S): MARPAT 143:13254

AB A contrast agent of formula I: V - L - R (I) where V is an organic group having binding affinity for an angiotensin II receptor site, L is a linear or branched amino acid-comprising biomodifier or linker moiety, and R is a reporter moiety detectable in in vivo ***imaging*** of a human or animal body. Contrast agents targeting the AT1 receptor may be suitable for detecting diseases such as congestive heart failure, atherosclerosis, and fibrosis.

L36 ANSWER 2 OF 25 HCAPLUS COPYRIGHT 2007 ACS on STN DUPLICATE 2
ACCESSION NUMBER: 2005:471932 HCAPLUS Full-text
DOCUMENT NUMBER: 143:26884
TITLE: Preparation of radiolabeled sulfonamide hydroxamate matrix metalloproteinase inhibitors as ***imaging*** agents
INVENTOR(S): ***Cuthbertson, Alan; Solbakken, Magne*** ; Bjurgert, Emma
PATENT ASSIGNEE(S): Amersham PLC, UK
SOURCE: PCT Int. Appl., 79 pp.
CODEN: PIXXD2
DOCUMENT TYPE: Patent
LANGUAGE: English
FAMILY ACC. NUM. COUNT: 1
PATENT INFORMATION:

| PATENT NO. | KIND | DATE | APPLICATION NO. | DATE |
|---|---|---|---|---|
| WO 2005049005 | A1 | 20050602 | WO 2004-GB4792 | 20041112 <-- |
| W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BW, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, EG, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NA, NI, | | | | |

NO, NZ, OM, PG, PH, PL, PT, RO, RU, SC, SD, SE, SG, SK, SL, SY,
TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, YU, ZA, ZM, ZW
RW: BW, GH, GM, KE, LS, MW, MZ, NA, SD, SL, SZ, TZ, UG, ZM, ZW, AM,
AZ, BY, KG, KZ, MD, RU, TJ, TM, AT, BE, BG, CH, CY, CZ, DE, DK,
EE, ES, FI, FR, GB, GR, HU, IE, IS, IT, LU, MC, NL, PL, PT, RO,
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NE, SN, TD, TG

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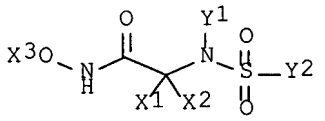
|---|---|---|---|---|---|
| AU 2004290950 | A1 | 20050602 | AU 2004-290950 | | 20041112 <-- |
| CA 2545267 | A1 | 20050602 | CA 2004-2545267 | | 20041112 <-- |
| EP 1682113 | A1 | 20060726 | EP 2004-798512 | | 20041112 <-- |
| R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, FI, RO, CY, TR, BG, CZ, EE, HU, PL, SK, IS | | | | | |
| BR 2004016528 | A | 20070109 | BR 2004-16528 | | 20041112 <-- |
| CN 1901894 | A | 20070124 | CN 2004-80040362 | | 20041112 <-- |
| NO 2006002118 | A | 20060511 | NO 2006-2118 | | 20060511 <-- |
| PRIORITY APPLN. INFO.: | | | GB 2003-26546 | A | 20031114 <-- |
| | | | WO 2004-GB4792 | W | 20041112 |

OTHER SOURCE(S): MARPAT 143:26884
GI



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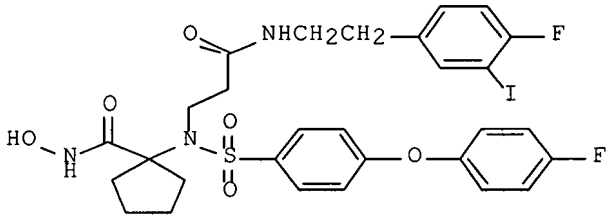

II

AB The present invention discloses that ***imaging*** agents I [Y1 = H, (CH2)nC(O)Z; n = 1-6; Z = OH, C1-6 alkoxy, C4-10 aryloxy, NR1R2; R1, R2 = independently H, C1-6 alkyl, C3-6 cycloalkyl, C1-6 fluoroalkyl, C4-10 aryl; X1 and X2 form C3-10 cycloalkyl or heterocyclic ring; X3 = H, C1-3 alkyl, C1-3 fluoroalkyl; Y2 = A1pOqA2; p, q = 0-1; A1 = C1-10 alkylene, C3-8 cycloalkylene, C1-10 perfluoroalkylene, C6-10 arylene, C2-10 heteroarylene; A2 = H, C1-10 alkyl, C3-8 cycloalkyl, C1-10 perfluoroalkyl, C6-10 aryl, C2-10 heteroaryl] which comprise a specific type of matrix metalloproteinase inhibitors (MMPi's) of the sulfonamide hydroxamate class labeled with an ***imaging*** moiety, are useful diagnostic ***imaging*** agents for in vivo ***imaging*** and diagnosis of the mammalian body. Thus, sulfonamides II (R = iodo) and related compds. were prepared and studies for MMP receptor binding specificity. II and its iodine-123 analog were also studied for biodistribution in a LLC tumor model in vivo, as well as in a ApoE ligation model.

RETABLE

| Referenced Author (RAU) | Year (RPY) | VOL (RVL) | PG (RPG) | Referenced Work (RWK) | Referenced File |
|---|---|---|---|---|---|
| Langley, K | 2002 | | | US 2002090654 A1 | HCAPLUS |

| | | | | |
|---|---|---|---|---|
| Mobashery, S | 2002 | | US 2002037916 A1 | HCAPLUS |
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| Storey, A | 2004 | | WO 2004069365 A | |

L36 ANSWER 3 OF 25 HCAPLUS COPYRIGHT 2007 ACS on STN DUPLICATE 3
ACCESSION NUMBER: 2005:426470 HCAPLUS Full-text
DOCUMENT NUMBER: 142:469186
TITLE: Conjugated angiotensin II analogs as ***imaging*** and therapeutic agents
INVENTOR(S): ***Cuthbertson, Alan***; Indrevoll, Bard; Eriksen, Morten
PATENT ASSIGNEE(S): Amersham Health A/S, Norway
SOURCE: PCT Int. Appl., 37 pp.
CODEN: PIXXD2
DOCUMENT TYPE: Patent
LANGUAGE: English
FAMILY ACC. NUM. COUNT: 3
PATENT INFORMATION:

| PATENT NO. | KIND | DATE | APPLICATION NO. | DATE |
|---|---|---|---|---|
| WO 2005044313 | A2 | 20050519 | WO 2004-NO335 | 20041105 <-- |
| WO 2005044313 | A3 | 20060511 | | |
| W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BW, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, EG, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NA, NI, NO, NZ, OM, PG, PH, PL, PT, RO, RU, SC, SD, SE, SG, SK, SL, SY, TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, YU, ZA, ZM, ZW | | | | |
| RW: BW, GH, GM, KE, LS, MW, MZ, NA, SD, SL, SZ, TZ, UG, ZM, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM, AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE, IS, IT, LU, MC, NL, PL, PT, RO, SE, SI, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG | | | | |
| EP 1699495 | A2 | 20060913 | EP 2004-800183 | 20041105 <-- |
| R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, LT, LV, FI, RO, MK, CY, AL, TR, BG, CZ, EE, HU, PL, SK, HR, IS, YU | | | | |
| CN 1901941 | A | 20070124 | CN 2004-80039758 | 20041105 <-- |
| PRIORITY APPLN. INFO.: | | | NO 2003-4952 A | 20031106 <-- |
| | | | GB 2004-16062 A | 20040719 |
| | | | WO 2004-NO335 W | 20041105 |

OTHER SOURCE(S): MARPAT 142:469186
AB The invention comprises pharmaceuticals of formula (I) Z-(L)n-V, wherein V denotes a peptide, L denotes an optional linker, Z denotes a group that optionally can carry an ***imaging*** moiety M, n denotes 0 or 1. The pharmaceuticals are active as therapeutic agents for the treatment of heart failure, cardiac arrhythmias and diseases where fibrosis is prominent such as COPD, liver fibrosis and atherosclerosis and are also useful as diagnostic agents for the diagnosis of heart failure and diseases were fibrosis is prominent such as COPD, liver fibrosis and atherosclerosis.

L36 ANSWER 4 OF 25 HCAPLUS COPYRIGHT 2007 ACS on STN DUPLICATE 4
ACCESSION NUMBER: 2005:120957 HCAPLUS Full-text
DOCUMENT NUMBER: 142:219561
TITLE: Preparation of peptide-based compounds as diagnostic ***imaging*** agents
INVENTOR(S): ***Cuthbertson, Alan; Solbakken, Magne***
PATENT ASSIGNEE(S): Amersham Health AS, Norway

SOURCE: PCT Int. Appl., 40 pp.
CODEN: PIXXD2
DOCUMENT TYPE: Patent
LANGUAGE: English
FAMILY ACC. NUM. COUNT: 1
PATENT INFORMATION:

| PATENT NO. | KIND | DATE | APPLICATION NO. | DATE |
|---|---|---|---|---|
| WO 2005012335 | A1 | 20050210 | WO 2004-GB3150 | 20040721 <-- |
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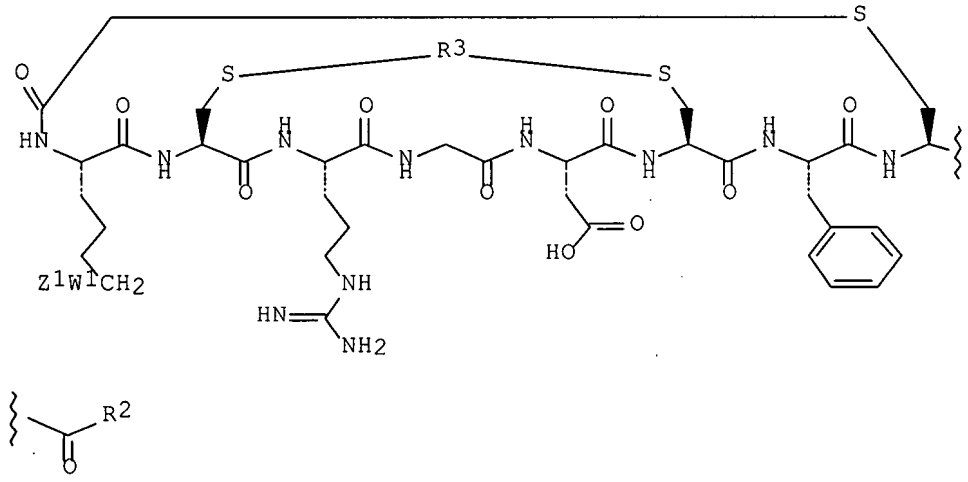
| RW: BW, GH, GM, KE, LS, MW, MZ, NA, SD, SL, SZ, TZ, UG, ZM, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM, AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE, IT, LU, MC, NL, PL, PT, RO, SE, SI, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG | | | | |
| CA 2533321 | A1 | 20050210 | CA 2004-2533321 | 20040721 <-- |
| EP 1648925 | A1 | 20060426 | EP 2004-743485 | 20040721 <-- |
| R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, FI, RO, CY, TR, BG, CZ, EE, HU, PL, SK | | | | |
| CN 1829735 | A | 20060906 | CN 2004-80022044 | 20040721 <-- |
| BR 2004012986 | A | 20061003 | BR 2004-12986 | 20040721 <-- |
| HU 200600230 | A2 | 20070129 | HU 2006-230 | 20040721 <-- |
| US 2006193773 | A1 | 20060831 | US 2006-566487 | 20060130 <-- |
| NO 2006000825 | A | 20060329 | NO 2006-825 | 20060221 <-- |

PRIORITY APPLN. INFO.: GB 2003-17815 A 20030730 <--
WO 2004-GB3150 W 20040721

OTHER SOURCE(S): CASREACT 142:219561; MARPAT 142:219561
GI







I

AB The invention relates to compds. I [R2 is [NH(CH2CH2O)3CH2CH2NHCOCH2OCH2CO ]0-10NH2; R3 is an alkylene or alkenylene bridge; W1 is absent or a spacer moiety (hetero)hydrocarbyl preferably derived from glutaric and/or succinic acid and/or a polyethylene glycol-based unit and/or a unit [NH(CH2CH2O)3CH2CH2NHCOCH2OCH2CO]n; Z1 is an antineoplastic agent, a chelating agent or a reporter moiety] and their use as targeting vectors that bind to receptors associated with angiogenesis. Compds. I may thus be used for diagnosis or therapy of various diseases. Thus, compound I [R2 is NH2, R3 is CH2, Z1-W1 is FCH2CH2SCH2CONH(CH2CH2O)5CH2CH2NHCOCH2OCH2CONH] was prepared via the solid-phase method and showed Ki = 7 nmol in an αvβ3 integrin receptor binding assay.

RETABLE

| Referenced Author (RAU) | Year (RPY) | VOL (RVL) | PG (RPG) | Referenced Work (RWK) | Referenced File |
|---|---|---|---|---|---|
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| Indrevoll, B | 2001 | | | WO 0177145 A | HCAPLUS |
| Indrevoll, B | 2003 | | | WO 03006491 A | HCAPLUS |
| Lister-James, J | 1999 | | | US 5888474 A | HCAPLUS |
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| Srinivasan, A | 2002 | | | WO 0220610 A | HCAPLUS |

L36 ANSWER 5 OF 25 HCAPLUS COPYRIGHT 2007 ACS on STN DUPLICATE 5
ACCESSION NUMBER: 2005:34777 HCAPLUS Full-text
DOCUMENT NUMBER: 142:130349
TITLE: Fluorescein-labeled peptides
INVENTOR(S): ***Cuthbertson, Alan***; Indrevoll, Bard; ***Solbakken, Magne***
PATENT ASSIGNEE(S): Amersham Health A/S, Norway
SOURCE: PCT Int. Appl., 43 pp.
CODEN: PIXXD2
DOCUMENT TYPE: Patent
LANGUAGE: English
FAMILY ACC. NUM. COUNT: 1
PATENT INFORMATION:

| PATENT NO. | KIND | DATE | APPLICATION NO. | | DATE |
|---|---|---|---|---|---|
| WO 2005003166 | A1 | 20050113 | WO 2004-NO208 | | 20040707 <-- |
| W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BW, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, EG, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NA, NI, NO, NZ, OM, PG, PH, PL, PT, RO, RU, SC, SD, SE, SG, SK, SL, SY, TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, YU, ZA, ZM, ZW | | | | | |
| RW: BW, GH, GM, KE, LS, MW, MZ, NA, SD, SL, SZ, TZ, UG, ZM, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM, AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE, IT, LU, MC, NL, PL, PT, RO, SE, SI, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG | | | | | |
| EP 1639004 | A1 | 20060329 | EP 2004-748784 | | 20040707 <-- |
| R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, FI, RO, CY, TR, BG, CZ, EE, HU, PL, SK | | | | | |
| CN 1820025 | A | 20060816 | CN 2004-80019515 | | 20040707 <-- |
| NO 2005006055 | A | 20060213 | NO 2005-6055 | | 20051219 <-- |
| PRIORITY APPLN. INFO.: | | | NO 2003-3115 | A | 20030708 <-- |
| | | | WO 2004-NO208 | W | 20040707 |

AB The invention relates to new peptide-based compds. and their use in diagnostic optical ***imaging***. More specifically the invention relates to the use of such peptide-based compds. as targeting vectors that bind to receptors associated with angiogenesis. The compds. are labeled with fluorescein and may be used as contrast agents in optical ***imaging*** in diagnosis of angiogenesis-related diseases.

RETABLE

| Referenced Author (RAU) | Year (RPY) | VOL (RVL) | PG (RPG) | Referenced Work (RWK) | Referenced File |
|---|---|---|---|---|---|
| Cuthbertson, A | 2002 | | | WO 0226776 A | HCAPLUS |
| de Groot, F | 2002 | 1 | 901 | MOLECULAR CANCER THE | HCAPLUS |
| Hellebust, H | 2002 | | | US 2002102217 A1 | HCAPLUS |
| Indrevoll, B | 2001 | | | WO 0177145 A | HCAPLUS |
| Katada, J | 1997 | 272 | 7720 | THE JOURNAL OF BIOLO | HCAPLUS |
| Riecke, B | 2001 | 33 | 307 | HORMONE AND METABOLI | HCAPLUS |
| Univ Leipzig | 1999 | | | DE 19808591 A | HCAPLUS |

L36 ANSWER 6 OF 25 HCAPLUS COPYRIGHT 2007 ACS on STN DUPLICATE 6
ACCESSION NUMBER: 2004:610037 HCAPLUS Full-text
DOCUMENT NUMBER: 141:145687
TITLE: Contrast agents for ***imaging*** angiotensin II receptors
INVENTOR(S): ***Solbakken, Magne***; Engell, Torgrim; Wadsworth, Harry John; Archer, Colin M.
PATENT ASSIGNEE(S): Amersham Health As, Norway
SOURCE: PCT Int. Appl., 31 pp.
CODEN: PIXXD2
DOCUMENT TYPE: Patent
LANGUAGE: English
FAMILY ACC. NUM. COUNT: 1
PATENT INFORMATION:

| PATENT NO. | KIND | DATE | APPLICATION NO. | | DATE |
|---|---|---|---|---|---|
| WO 2004062568 | A2 | 20040729 | WO 2004-NO2 | | 20040109 <-- |
| WO 2004062568 | A3 | 20040930 | | | |
| W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BW, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, EG, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ | | | | | |
| EP 1581262 | A2 | 20051005 | EP 2004-701136 | | 20040109 <-- |
| R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, LT, LV, FI, RO, MK, CY, AL, TR, BG, CZ, EE, HU, SK | | | | | |
| CN 1723043 | A | 20060118 | CN 2004-80001954 | | 20040109 <-- |
| JP 2006517539 | T | 20060727 | JP 2006-500737 | | 20040109 <-- |
| IN 2005DN02527 | A | 20061229 | IN 2005-DN2527 | | 20050611 <-- |
| US 2007020176 | A1 | 20070125 | US 2006-541949 | | 20060607 <-- |
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| | | | WO 2004-NO2 | W | 20040109 |

OTHER SOURCE(S): MARPAT 141:145687
AB The present invention relates to contrast agents in which the targeting vector binds to angiotensin II receptors. The targeting vector, a receptor antagonist such as losartan, valsartan, candesartan or eprosartan, is conjugated via a spacer or linker to a moiety detectable in in vivo ***imaging*** procedures. The ***imaging*** moiety is a chelated radionuclide such as 99mTc. The ***imaging*** moiety may also consist of paramagnetic or fluorescent metal ions or other detectable species.

L36 ANSWER 7 OF 25 HCAPLUS COPYRIGHT 2007 ACS on STN DUPLICATE 7
ACCESSION NUMBER: 2003:58113 HCAPLUS Full-text
DOCUMENT NUMBER: 138:122862
TITLE: Preparation of peptide-based compounds as diagnostic *imaging* agents
INVENTOR(S): ***Cuthbertson, Alan***; Indrevoll, Bard; ***Solbakken, Magne***
PATENT ASSIGNEE(S): Amersham Health AS, Norway
SOURCE: PCT Int. Appl., 56 pp.
CODEN: PIXXD2
DOCUMENT TYPE: Patent
LANGUAGE: English
FAMILY ACC. NUM. COUNT: 2
PATENT INFORMATION:

| PATENT NO. | KIND | DATE | APPLICATION NO. | DATE |
|---|---|---|---|---|
| WO 2003006491 | A2 | 20030123 | WO 2002-NO250 | 20020708 <-- |
| WO 2003006491 | A3 | 20031224 | | |
| W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, OM, PH, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VN, YU, ZA, ZM, ZW | | | | |
| RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM, AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG | | | | |
| CA 2452923 | A1 | 20030123 | CA 2002-2452923 | 20020708 <-- |
| AU 2002313616 | A1 | 20030129 | AU 2002-313616 | 20020708 <-- |
| EP 1404371 | A2 | 20040407 | EP 2002-753307 | 20020708 <-- |
| R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, MC, PT, IE, SI, LT, LV, FI, RO, MK, CY, AL, BG, CZ, EE | | | | |
| BR 2002010886 | A | 20040622 | BR 2002-10886 | 20020708 <-- |
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| JP 2005507376 | T | 20050317 | JP 2003-512261 | 20020708 <-- |
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| US 2005070466 | A1 | 20050331 | US 2004-753729 | 20040108 <-- |
| NO 2004000084 | A | 20040309 | NO 2004-84 | 20040109 <-- |
| PRIORITY APPLN. INFO.: | | | GB 2001-16815 A | 20010710 <-- |
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| | | | WO 2002-NO250 W | 20020708 <-- |

OTHER SOURCE(S): MARPAT 138:122862
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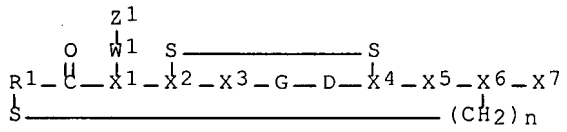

I

AB Peptide-based compds. I [G represents glycine; D represents aspartic acid; R1 = (CH2)1-10 or (CH2)1-10C6H4, n = 1 or 2; X1 represents an amino acid residue which possesses a functional side chain such as an acid or amine; X2, X4 represent an amino acid residue capable of forming a disulfide bond; X3

represents arginine, N-methylarginine or an arginine mimetic; X5 represents a hydrophobic amino acid or derivative; X6 represents a thiol-containing amino acid residue; X7 is absent or represents a biomodifier moiety; Z1 represents an antineoplastic agent, a chelating agent or a reporter moiety; W1 is absent or represents a spacer moiety] or pharmaceutically-acceptable salts were prepared for use as diagnostic ***imaging*** agents or as therapeutic agents which comprise targeting vectors which bind to integrin receptors. Thus, cyclo[CH2CO-Lys(cPn216- glutaryl)-Cys2-Arg-Gly-Asp-Cys6-Phe-Cys]-NH2 disulfide (Cys2-6) [cPn216 is technetium chelate residue (HON:CMeCMe2CH2CH2)2CHCH2CH2NH] was prepared via the solid-phase method.

L36 ANSWER 8 OF 25 HCAPLUS COPYRIGHT 2007 ACS on STN DUPLICATE 8
ACCESSION NUMBER: 2003:242904 HCAPLUS Full-text
DOCUMENT NUMBER: 139:328246
TITLE: Amphiphilic lipopeptide microparticles as contrast agents for medical ultrasound ***imaging***
AUTHOR(S): ***Cuthbertson, Alan***; Tornes, Audun; ***Solbakken, Magne***; Moen, Ove; Eriksen, Morten
CORPORATE SOURCE: Dep. of Exploratory Res., Amersham Health AS, Oslo, Norway
SOURCE: Macromolecular Bioscience (***2003***), 3(1), 11-17
CODEN: MBAIBU; ISSN: 1616-5187
PUBLISHER: Wiley-VCH Verlag GmbH & Co. KGaA
DOCUMENT TYPE: Journal
LANGUAGE: English
AB In this study the authors investigated the utility of complementary amphiphilic lipopeptides as a new membrane formulation suitable for the preparation of gas-filled microbubbles. Through primarily ion pairing and hydrophobic interactions we rationalized that the stacking of synthetic lipopeptides into the surface of microbubbles would make bubble suspensions useful as ultrasound contrast agents. By mixing charged lipopeptides in propylene glycol/glycerol solns. in the presence of a perfluorocarbon gas followed by vigorous shaking, microbubble suspensions were formed in good yield with a size distribution spanning the range 1-7+10-6 m. The microbubbles were studied in an in vivo model and provided ***imaging*** efficacy comparable with conventional ultrasound contrast agents.

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L36 ANSWER 9 OF 25 HCAPLUS COPYRIGHT 2007 ACS on STN DUPLICATE 9
ACCESSION NUMBER: 2002:695821 HCAPLUS Full-text
DOCUMENT NUMBER: 137:237702
TITLE: Improved peptide-chelate conjugates
INVENTOR(S): ***Cuthbertson, Alan***; Mendizabal, Marivi; Dixon, Mark; Storey, Anthony Eamon
PATENT ASSIGNEE(S): Amersham PLC, UK
SOURCE: PCT Int. Appl., 50 pp.
CODEN: PIXXD2
DOCUMENT TYPE: Patent
LANGUAGE: English
FAMILY ACC. NUM. COUNT: 1
PATENT INFORMATION:

| PATENT NO. | KIND | DATE | APPLICATION NO. | DATE |
|---|---|---|---|---|
| WO 2002070018 | A2 | 20020912 | WO 2002-GB857 | 20020301 <-- |
| WO 2002070018 | A3 | 20021205 | | |
| W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, OM, PH, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VN, YU, ZA, ZM, ZW | | | | |
| RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG | | | | |
| CA 2439579 | A1 | 20020912 | CA 2002-2439579 | 20020301 <-- |
| EP 1368064 | A2 | 20031210 | EP 2002-701431 | 20020301 <-- |
| R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, LT, LV, FI, RO, MK, CY, AL, TR | | | | |
| JP 2004524323 | T | 20040812 | JP 2002-569189 | 20020301 <-- |
| US 2006222593 | A1 | 20061005 | US 2004-469801 | 20041210 <-- |
| PRIORITY APPLN. INFO.: | | | GB 2001-5224 A | 20010302 <-- |
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OTHER SOURCE(S): MARPAT 137:237702
AB A peptide-chelate conjugate with affinity for the ST receptor is disclosed, wherein the chelate is tetradentate. The peptide-chelate conjugate of the invention may be labeled with a radiometal to provide a metal complex. A radiopharmaceutical composition comprising the metal complex is provided, which is suitable for the diagnostic ***imaging*** of colorectal cancer. Also provided for in the invention is a kit for the preparation of the radiopharmaceutical preparation

L36 ANSWER 10 OF 25 HCAPLUS COPYRIGHT 2007 ACS on STN DUPLICATE 10
ACCESSION NUMBER: 2002:256296 HCAPLUS Full-text
DOCUMENT NUMBER: 136:263481
TITLE: Preparation of peptide-based compounds as diagnostic ***imaging*** agents
INVENTOR(S): ***Cuthbertson, Alan***
PATENT ASSIGNEE(S): Nycomed Imaging AS, Norway; Amersham Health AS
SOURCE: PCT Int. Appl., 39 pp.
CODEN: PIXXD2
DOCUMENT TYPE: Patent
LANGUAGE: English
FAMILY ACC. NUM. COUNT: 1
PATENT INFORMATION:

| PATENT NO. | KIND | DATE | APPLICATION NO. | DATE |
|---|---|---|---|---|

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|---|---|---|---|---|
| WO 2002026776 | A2 | 20020404 | WO 2001-NO390 | 20010925 <-- |
| WO 2002026776 | A3 | 20030828 | | |
| W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, PH, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, US, UZ, VN, YU, ZA, ZW | | | | |
| RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG | | | | |
| CA 2420577 | A1 | 20020404 | CA 2001-2420577 | 20010925 <-- |
| AU 2001092456 | A5 | 20020408 | AU 2001-92456 | 20010925 <-- |
| EP 1358206 | A2 | 20031105 | EP 2001-972816 | 20010925 <-- |
| R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, LT, LV, FI, RO, MK, CY, AL, TR | | | | |
| JP 2004509975 | T | 20040402 | JP 2002-531159 | 20010925 <-- |
| NO 2003001323 | A | 20030324 | NO 2003-1323 | 20030324 <-- |
| US 2003176639 | A1 | 20030918 | US 2003-395500 | 20030324 <-- |
| PRIORITY APPLN. INFO.: | | | NO 2000-4795 A | 20000926 <-- |
| | | | US 2001-259919P P | 20010105 <-- |
| | | | WO 2001-NO390 W | 20010925 <-- |

OTHER SOURCE(S): MARPAT 136:263481

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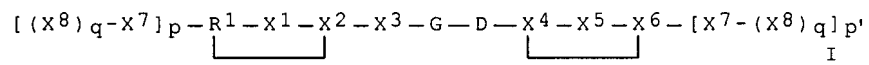


AB Peptide-based compds. I [G represents glycine; D represents aspartic acid; p, p' = 0, 1 and p + p' = 1; when p = 0 then R1 is (CH2)nCO or (CH2)nC6H4CO, where n = 1-5, when p = 1 then R1 is or one or more bridge-forming amino acids; X1 = a bond or 1-5 amino acids, an amino acid derivatized with a carbohydrate moiety, an amino acid functionalized with a spacer or linker and/or a chelate binding or capable of binding a reporter suitable for in vivo ***imaging***; X2, X4 are cysteine, homocysteine or other amino acids capable of forming a cyclizing bond such as aspartic acid and lysine; X3 is arginine, N-methylarginine, or an arginine mimetic; X5 is a hydrophobic amino acid; X6 is an amino acid capable of forming a cyclizing bond; X7 is a bond or 1-10 amino acids or a spacer or linker, optionally allowing for labeling with multiple chelates as defined by X8, and optionally comprising one or more ethylene glycol units or any other spacer component; X8 is a chelate binding to, or capable of binding, a metal radionuclide or any other reporter suitable for in vivo ***imaging***, NH2 or is absent; q is 0-8; one of the bridges (between R1 and X2 or between X4 and X6) comprises a disulfide bond] were prepared for use in therapeutically effective treatments and as diagnostic ***imaging*** agents. More specifically, the invention relates to the use of such peptide-based compds. used as targeting vectors that bind to receptors associated with angiogenesis, in particular the αvβ3 integrin receptor. Synthesis and conjugation of peptide vector H-Ala-Cys-Asp-Cys-Arg-Gly-Asp-Cys-Phe-Cys-Gly-OH with disulfide bonds connecting Cys-2 and Cys-4 and Cys-8 and Cys-10 and technetium chelate-succinic acid intermediate [HON:CMeCMe2NHCH2CH2]2NCH2CH2NHCOCH2CH2 CO2H are described.

L36 ANSWER 11 OF 25 HCAPLUS COPYRIGHT 2007 ACS on STN DUPLICATE 11
ACCESSION NUMBER: 2002:386498 HCAPLUS Full-text
DOCUMENT NUMBER: 138:52011
TITLE: In vivo ***imaging*** of human colon cancer xenografts in immunodeficient mice using a guanylyl cyclase C-specific ligand
AUTHOR(S): Wolfe, Henry R.; Mendizabal, Marivi; Lleong, Elinor; ***Cuthbertson, Alan***; Desai, Vinay; Pullan, Shirley; Fujii, Dennis K.; Morrison, Matthew; Pither, Richard; Waldman, Scott A.
CORPORATE SOURCE: Research and Development Department, Targeted Diagnostics and Therapeutics, Inc., West Chester, PA, 19380, USA
SOURCE: Journal of Nuclear Medicine (***2002***), 43(3), 392-399
CODEN: JNMEAQ; ISSN: 0161-5505
PUBLISHER: Society of Nuclear Medicine
DOCUMENT TYPE: Journal
LANGUAGE: English

AB Guanylyl cyclase C (GC-C) is a transmembrane receptor expressed by human intestinal cells and primary and metastatic colorectal adenocarcinomas but not by extraintestinal tissues or tumors. The Escherichia coli heat-stable enterotoxin analog, STa (5-18), is a 14-amino acid peptide that selectively binds to the extracellular domain of GC-C with subnanomolar affinity. This study examined the utility of a radiolabeled conjugate of STa (5-18) to selectively target and image extraintestinal human colon cancer xenografts in vivo in nude mice. The STa conjugate, ethoxyethyl-mercaptoacetamidoadipoylglycylglycine-STa (5-18) (NC100586), was synthesized and labeled with 99mTc to produce 99mTc-NC100586. This compound was i.v. administered to nude mice bearing human colon cancer xenografts, and specific targeting was evaluated by biodistribution and gamma camera ***imaging***. In CD-1 nude mice, biodistribution and scintigraphic ***imaging*** analyses showed selective uptake of 99mTc-NC100586 into human colon cancer xenografts that express GC-C but not into normal tissues that do not express GC-C. Similarly, 99mTc-NC100586 injected i.v. into CD-1 nude mice with human colon cancer hepatic metastases selectively accumulated in those metastases, and .apprx.5-mm foci of tumor cells were visualized after ex vivo ***imaging*** of excised livers. Accumulation of 99mTc-NC100586 in human colon cancer xenografts reflected binding to GC-C because 99mTc-NC100588, an inactive analog that does not bind to GC-C, did not selectively accumulate in cancer xenografts compared with normal tissues. Also, coadministration of excess unlabeled STa (5-18) prevented accumulation of 99mTc-NC100586 in human colon cancer xenografts. Furthermore, 99mTc-NC100586 did not selectively accumulate in Lewis lung tumor xenografts, which do not express GC-C. This study showed that i.v. administered STa (5-18) selectively recognizes and binds to GC-C expressed by human colon cancer cells in vivo. Also shown was the ability to exploit this selective interaction to target ***imaging*** agents to extraintestinal human colon tumors in nude mice. These results suggest the utility of STa and GC-C for the development of novel targeted ***imaging*** and therapeutic agents with high specificity for metastatic colorectal tumors in humans.

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L36 ANSWER 12 OF 25 HCAPLUS COPYRIGHT 2007 ACS on STN DUPLICATE 12
ACCESSION NUMBER: 2001:763033 HCAPLUS Full-text
DOCUMENT NUMBER: 135:318716
TITLE: Preparation of peptide-based compounds as diagnostic ***imaging*** agents
INVENTOR(S): ***Cuthbertson, Alan***; Indrevoll, Bard
PATENT ASSIGNEE(S): Nycomed Imaging AS, Norway
SOURCE: PCT Int. Appl., 63 pp.
CODEN: PIXXD2
DOCUMENT TYPE: Patent
LANGUAGE: English
FAMILY ACC. NUM. COUNT: 1
PATENT INFORMATION:

| PATENT NO. | KIND | DATE | APPLICATION NO. | DATE |
|---|---|---|---|---|
| WO 2001077145 | A2 | 20011018 | WO 2001-NO146 | 20010406 <-- |
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RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG

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| EP 1272507 | B1 | 20050629 | | | |
| R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, LT, LV, FI, RO, MK, CY, AL, TR | | | | | |
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| ES 2244607 | T3 | 20051216 | ES 2001-1924011 | | 20010406 <-- |
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| PRIORITY APPLN. INFO.: | | | GB 2000-9042 | A | 20000412 <-- |
| | | | US 2000-211337P | P | 20000613 <-- |
| | | | GB 2000-25070 | A | 20001012 <-- |
| | | | WO 2001-NO146 | W | 20010406 <-- |

OTHER SOURCE(S): MARPAT 135:318716

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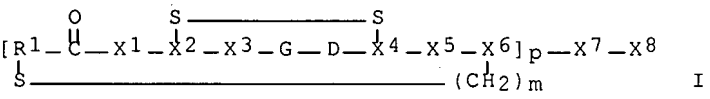

I

AB Peptide-based compds. I [G represents glycine; D represents aspartic acid; R1 = (CH2)n or (CH2)nC6H4 (n = 1-10), m = 1 or 2; p = 1-10; X1 represents a bond or 1-5 amino acid residues which can independently be derivatized with a functional side chain suitable for modifying pharmacokinetics or blood clearance rates and can bind a reporter (R) moiety suitable for in vivo ***imaging*** via a linker (L) moiety, a chelating agent or an L moiety attached to a chelating agent; X2, X4 represent an amino acid residue capable of forming a disulfide bond; X3 represents arginine, N-methylarginine or an arginine mimetic; X5 represents a hydrophobic amino acid or derivative; X6 represents a thiol-containing amino acid residue; X7 represents an L moiety or 1-10 amino acid residues, optionally as part of an L moiety, with the properties of X1; X7 is absent; X8 represents an R moiety or NH2 or is absent] or pharmaceutically acceptable salts were prepared for use as diagnostic ***imaging*** agents or as therapeutic agents which comprise a targeting vector which binds to receptors associated with integrin receptors. Thus, [Cys2-6]cyclo[CH2CONH-Asp-Cys-Arg-Gly-Asp- Cys-Phe-Cys]-Gly-NH(CH2CH2O)2CH2CH2NHCO(CH2)3CH(NHCOCH2SCHMeOEt)CO-Gly-Gly- OH (VIa) was prepared by solid-phase peptide coupling, cyclization, deprotection, and conjugation with N3S-adipate chelator active ester. Compd.VIa was labeled with technetium.

L36 ANSWER 13 OF 25 HCAPLUS COPYRIGHT 2007 ACS on STN DUPLICATE 13

| | |
|---|---|
| ACCESSION NUMBER: | 2001:519335 HCAPLUS Full-text |
| DOCUMENT NUMBER: | 135:111977 |
| TITLE: | Diagnostic/therapeutic agents having phospholipid-based microbubbles coupled to one or more vectors |
| INVENTOR(S): | Klaveness, Jo; Rongved, Pal; Hogset, Anders; Tolleshaug, Helge; Naevestad, Anne; Hellebust, |

Halldis; Hoff, Lars; ***Cuthbertson, Alan;*** Lovhaug, Dagfinn; ***Solbakken, Magne***
PATENT ASSIGNEE(S): Nycomed Imaging As, Norway
SOURCE: U.S., 89 pp., Cont.-in-part of U.S. Ser. No. 958,993.
CODEN: USXXAM

DOCUMENT TYPE: Patent
LANGUAGE: English
FAMILY ACC. NUM. COUNT: 10
PATENT INFORMATION:

| PATENT NO. | KIND | DATE | APPLICATION NO. | DATE |
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| US 6261537 | B1 | 20010717 | US 1997-960054 | 19971029 <-- |
| CN 1234742 | A | 19991110 | CN 1997-199047 | 19971028 <-- |
| HU 9904595 | A2 | 20000428 | HU 1999-4595 | 19971028 <-- |
| US 6331289 | B1 | 20011218 | US 1997-959206 | 19971028 <-- |
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| R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, FI, CY | | | | |
| ES 2224379 | T3 | 20050301 | ES 1998-917461 | 19980424 <-- |
| KR 2000052829 | A | 20000825 | KR 1999-703658 | 19990427 <-- |
| US 2002102215 | A1 | 20020801 | US 2001-765614 | 20010122 <-- |
| US 2002102217 | A1 | 20020801 | US 2001-925715 | 20010810 <-- |
| US 6680047 | B2 | 20040120 | | |
| CN 1440816 | A | 20030910 | CN 2002-160420 | 20021230 <-- |
| US 2004141922 | A1 | 20040722 | US 2003-722075 | 20031126 <-- |
| US 2005002865 | A1 | 20050106 | US 2003-734730 | 20031215 <-- |
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PRIORITY APPLN. INFO.:

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| GB 1997-699 | A | 19970115 <-- |
| GB 1997-8265 | A | 19970424 <-- |
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| GB 1997-11846 | A | 19970606 <-- |
| US 1997-49264P | P | 19970606 <-- |
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| US 1997-49268P | P | 19970606 <-- |
| US 1997-958993 | A2 | 19971028 <-- |
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| US 1997-960054 | A1 | 19971029 <-- |
| EP 1998-917461 | A3 | 19980424 <-- |
| US 2001-765614 | B1 | 20010122 <-- |
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| US 2003-722075 | A2 | 20031126 <-- |

AB Targetable diagnostic and/or therapeutically active agents, e.g. ultrasound contrast agents, having reporters comprise gas-filled microbubbles stabilized by monolayers of film-forming surfactants, the reporter being coupled or linked to at least one vector. The gas is air, nitrogen, oxygen, carbon dioxide, hydrogen, an inert gas, a sulfur fluoride, selenium hexafluoride, a low mol. weight hydrocarbon, a ketone, an ester, a halogenated low mol. weight hydrocarbon or their mixts. The film-forming surfactant material is one or more phospholipids selected from the group consisting of phosphatidylserines,

phosphatidylglycerols, phosphatidylinositols, phosphatidic acids and cardiolipins. A therapeutic agent is an antineoplastic agent, blood product, biol. response modifier, antifungal agent, hormone or hormone analog, vitamin, enzyme, antiallergic agent, tissue factor inhibitor, platelet inhibitor, coagulation protein target inhibitor, fibrin formation inhibitor, fibrinolysis promoter, antiangiogenic, circulatory drug, metabolic potentiator, antitubercular, antiviral, vasodilator, antibiotic, anti-inflammatory, antiprotozoal, antirheumatic, narcotic, opiate, cardiac glycoside, neuromuscular blocker, sedative, local anesthetic, general anesthetic or genetic material. For example, an endothelial cell adhesion of phosphatidylserine-encapsulated perfluorobutane microbubbles coated with polylysine was higher than adhesion of uncoated microbubbles. Also, a thrombus was detected by ultrasound in patients with suspected venous thrombosis using i.v. phosphatidylserine-encapsulated microbubbles. The microbubbles contained inactivated human thrombin-succinyl-PEG 3400-distearoylphosphatidylethanol amine incorporated into the encapsulating membrane.

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| Unger | 1998 | | | US 5846517 | HCAPLUS |
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L36 ANSWER 14 OF 25 HCAPLUS COPYRIGHT 2007 ACS on STN DUPLICATE 14
ACCESSION NUMBER: 2000:253997 HCAPLUS Full-text
DOCUMENT NUMBER: 132:284295
TITLE: Contrast agents
INVENTOR(S): Klaveness, Jo; Naevestad, Anne; ***Cuthbertson, Alan; Solbakken, Magne***
PATENT ASSIGNEE(S): Nycomed Imaging As, Norway
SOURCE: U.S., 31 pp., Cont.-in-part of PCT 9818497.
CODEN: USXXAM
DOCUMENT TYPE: Patent
LANGUAGE: English
FAMILY ACC. NUM. COUNT: 10
PATENT INFORMATION:

| PATENT NO. | KIND | DATE | APPLICATION NO. | DATE |
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| US 6331289 | B1 | 20011218 | US 1997-959206 | 19971028 <-- |
| ES 2206689 | T3 | 20040516 | ES 1997-910517 | 19971028 <-- |
| US 2002102217 | A1 | 20020801 | US 2001-925715 | 20010810 <-- |
| US 6680047 | B2 | 20040120 | | |
| US 2005002865 | A1 | 20050106 | US 2003-734730 | 20031215 <-- |

PRIORITY APPLN. INFO.:

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| GB 1996-22364 | A | 19961028 <-- |
| GB 1996-22365 | A | 19961028 <-- |
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| GB 1997-9088 | A | 19970502 <-- |
| US 1997-48054P | P | 19970530 <-- |
| WO 1997-GB2657 | A2 | 19971028 <-- |
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| US 1997-49263P | P | 19970607 <-- |
| US 1997-49266P | P | 19970607 <-- |
| US 1997-959206 | A | 19971028 <-- |
| US 2001-925715 | A1 | 20010810 <-- |

OTHER SOURCE(S): MARPAT 132:284295
AB The invention provides a composition containing compds. with a nonpeptide organic group having binding affinity for an endothelin receptor site, a linker moiety or a bond, and a moiety detectable in in vivo ***imaging*** of a human or animal body. This composition of matter may be used to image diseases and disorders, particularly of the cardiovascular system. A compound was prepared from lysine and 27-O-3-[2-(3-carboxyacryloylamino)-5-

hydroxyphenyl]acryloyloxymycerone and the resulting compound treated with DTPA dianhydride to give a compound which was chelated with Gd or 99mTc.

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| Anon | 1996 | | | | HCAPLUS |
| Anon | 1996 | | | DE 19503644 | |
| Anon | 1996 | | | CA 2211364 | HCAPLUS |
| Anon | 1997 | | | DE 19536781 | HCAPLUS |
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| Us National Library Of | | | | Database Medline | |

L36 ANSWER 15 OF 25 HCAPLUS COPYRIGHT 2007 ACS on STN DUPLICATE 15
ACCESSION NUMBER: 1999:708651 HCAPLUS Full-text
DOCUMENT NUMBER: 131:319900
TITLE: Diagnostic/therapeutic agents comprising membrane-forming amphiphilic lipopeptide-stabilized gas microbubbles
INVENTOR(S): ***Cuthbertson, Alan; Solbakken, Magne*** ; Wolfe, Henry Raphael
PATENT ASSIGNEE(S): Marsden, John Christopher, UK; Nycomed Imaging A/S
SOURCE: PCT Int. Appl., 59 pp.
CODEN: PIXXD2
DOCUMENT TYPE: Patent
LANGUAGE: English
FAMILY ACC. NUM. COUNT: 1
PATENT INFORMATION:

| PATENT NO. | KIND | DATE | APPLICATION NO. | | DATE |
|---|---|---|---|---|---|
| WO 9955383 | A2 | 19991104 | WO 1999-GB1247 | | 19990422 <-- |
| WO 9955383 | A3 | 20000706 | | | |
| W: AE, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN, CU, CZ, DE, DK, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MD, MG, MK, MN, MX, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, UA, UG, US, UZ, VN, YU, ZA | | | | | |
| RW: GH, GM, KE, LS, MW, SD, SL, SZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG | | | | | |
| CA 2329778 | A1 | 19991104 | CA 1999-2329778 | | 19990422 <-- |
| EP 1073475 | A2 | 20010207 | EP 1999-918154 | | 19990422 <-- |
| R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, FI | | | | | |
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| ZA 2000005952 | A | 20010507 | ZA 2000-5952 | | 20001024 <-- |
| US 6548048 | B1 | 20030415 | US 2000-695273 | | 20001025 <-- |
| NO 2000005382 | A | 20001218 | NO 2000-5382 | | 20001026 <-- |
| PRIORITY APPLN. INFO.: | | | GB 1998-9084 | A | 19980428 <-- |
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AB Novel membrane-forming amphiphilic lipopeptides comprise one or more peptide moieties containing 2-50 aminoacyl residues and one or more hydrocarbon chains containing 5-50 carbon atoms. Such lipopeptides may be used in the formation of stabilized gas microbubble dispersions suitable for use as diagnostic and/or therapeutic agents, for example as ultrasound contrast agents. Perfluorobutane-containing microbubbles were prepared that used N-[3-(2-aminoethanamido)-5-[2-(n-hexadecyl)octadecanamido]benzoyl]gly cine (preparation given) as the membrane-forming agent.

L36 ANSWER 16 OF 25 HCAPLUS COPYRIGHT 2007 ACS on STN DUPLICATE 16
ACCESSION NUMBER: 1999:690991 HCAPLUS Full-text
DOCUMENT NUMBER: 131:308623
TITLE: Ultrasound ***imaging*** contrast agents, particularly for perfusion in the myocardium
INVENTOR(S): Eriksen, Morten; Tolleshaug, Helge; Skurtveit, Roald; ***Cuthbertson, Alan***; Ostensen, Jonny; Frigstad, Sigmund; Rongved, Pal
PATENT ASSIGNEE(S): Marsden, John Christopher, UK; Nycomed Imaging AS
SOURCE: PCT Int. Appl., 80 pp.
CODEN: PIXXD2
DOCUMENT TYPE: Patent
LANGUAGE: English
FAMILY ACC. NUM. COUNT: 1
PATENT INFORMATION:

| PATENT NO. | KIND | DATE | APPLICATION NO. | DATE |
|---|---|---|---|---|
| WO 9953963 | A1 | 19991028 | WO 1999-GB1221 | 19990422 <-- |
| W: AE, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN, CU, CZ, DE, DK, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MD, MG, MK, MN, MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, UA, UG, US, UZ, VN, YU, ZA, ZW | | | | |
| RW: GH, GM, KE, LS, MW, SD, SL, SZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG | | | | |
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| AU 9936172 | A | 19991108 | AU 1999-36172 | 19990422 <-- |
| BR 9909822 | A | 20001219 | BR 1999-9822 | 19990422 <-- |
| EP 1073473 | A1 | 20010207 | EP 1999-918133 | 19990422 <-- |
| R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, FI | | | | |
| HU 200102878 | A2 | 20020328 | HU 2001-2878 | 19990422 <-- |
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| ZA 2000005789 | A | 20010730 | ZA 2000-5789 | 20001018 <-- |
| NO 2000005250 | A | 20001218 | NO 2000-5250 | 20001019 <-- |
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| PRIORITY APPLN. INFO.: | | | GB 1998-8599 A | 19980422 <-- |
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AB Ultrasonic visualization of a subject, particularly of perfusion in the myocardium and other tissues, is performed using novel gas-containing contrast agent prepns. which promote controllable and temporary growth of the gas phase in vivo following administration and can therefore act as deposited perfusion tracers. The prepns. comprise an injectable aqueous medium comprising dispersed gas and an injectable oil-in-water emulsion in which the oil phase comprises a diffusible component capable of diffusion in vivo into the

dispersed gas to promote temporary growth thereof, such that material present at the surfaces of the dispersed gas phase and material present at the surfaces of the dispersed oil phase have affinity for each other, e.g. as a result of having opposite charges. In cardiac perfusion ***imaging*** the prepns. may advantageously be coadministered with vasodilator drugs such as adenosine in order to enhance the differences between return signal intensity from normal and hypoperfused myocardial tissue resp. A neg.-charged perfluorobutane gas dispersion and a pos.-charged perfluorodimethylcyclobutane emulsion were simultaneously injected i.v. into a dog. The resulting myocardial contrast effect was far more intense than that observed when the dispersion and emulsion were both neg.-charged. The contrast lasted for 20 min.

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L36 ANSWER 17 OF 25 HCAPLUS COPYRIGHT 2007 ACS on STN DUPLICATE 17
ACCESSION NUMBER: 1998:300865 HCAPLUS Full-text
DOCUMENT NUMBER: 129:4871
TITLE: Preparation of targetable diagnostic and therapeutic gas-containing or gas-generating ultrasound contrast agents
INVENTOR(S): Klaveness, Jo; Rongved, Pal; Hogset, Anders; Tolleshaug, Helge; ***Cuthbertson, Alan***; et al.
PATENT ASSIGNEE(S): Marsden, John Christopher, UK; Nycomed Imaging AS
SOURCE: PCT Int. Appl., 150 pp.
CODEN: PIXXD2
DOCUMENT TYPE: Patent
LANGUAGE: English
FAMILY ACC. NUM. COUNT: 10
PATENT INFORMATION:

| PATENT NO. | KIND | DATE | APPLICATION NO. | DATE |
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| RW: GH, KE, LS, MW, SD, SZ, UG, ZW, AT, BE, CH, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG, CI, CM, GA, GN, ML, MR, NE, SN, TD, TG | | | | |
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| MX 9903867 | A | 20000531 | MX 1999-3867 | | 19990426 <-- |
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| US 2002102217 | A1 | 20020801 | US 2001-925715 | | 20010810 <-- |
| US 6680047 | B2 | 20040120 | | | |
| US 2005002865 | A1 | 20050106 | US 2003-734730 | | 20031215 <-- |
| PRIORITY APPLN. INFO.: | | | GB 1996-22366 | A | 19961028 <-- |
| | | | GB 1996-22369 | A | 19961028 <-- |
| | | | GB 1997-2195 | A | 19970204 <-- |
| | | | GB 1997-8265 | A | 19970424 <-- |
| | | | GB 1997-11837 | A | 19970606 <-- |
| | | | GB 1997-11839 | A | 19970606 <-- |
| | | | US 1996-49263P | P | 19970606 <-- |
| | | | US 1996-49264P | P | 19970606 <-- |
| | | | US 1996-49266P | P | 19970606 <-- |
| | | | US 1997-49264P | P | 19970606 <-- |
| | | | US 1997-49263P | P | 19970607 <-- |
| | | | US 1997-49266P | P | 19970607 <-- |
| | | | US 1997-959206 | A | 19971028 <-- |
| | | | WO 1997-GB2953 | W | 19971028 <-- |
| | | | EP 1998-917461 | A3 | 19980424 <-- |
| | | | US 2001-925715 | A1 | 20010810 <-- |

AB Targetable diagnostic and/or therapeutically active agents, e.g. ultrasound contrast agents, comprising a suspension in an aqueous carrier liquid of a reporter comprising gas-containing or gas-generated material, in which the reporter is coupled or linked to one or more non-bioactive vectors. Thus, lipopeptide R-Lys(R)-Lys-Arg-Lys-Arg-Trp-Glu-Pro-Pro-Arg-Ala-Arg-Ile- OH (I; R = hexadecanoyl) (preparation given) containing a heparin binding site and a fibronectin binding site, was prepared by standard solid-phase methods. Microbubbles containing lipopeptide I were tested in vitro for binding to endothelial cells under flow conditions.

L36 ANSWER 18 OF 25 HCAPLUS COPYRIGHT 2007 ACS on STN DUPLICATE 18

ACCESSION NUMBER: 1998:300864 HCAPLUS Full-text

DOCUMENT NUMBER: 129:4870

TITLE: Preparation of targetable diagnostic and therapeutic ultrasound contrast agents

INVENTOR(S): Klaveness, Jo; Rongved, Pal; Hogset, Anders; Tolleshaug, Helge; Godal, Aslak; ***Cuthbertson, Alan***; et al.

PATENT ASSIGNEE(S): Marsdan, John Christopher, UK; Nycomed Imaging AS

SOURCE: PCT Int. Appl., 140 pp.
CODEN: PIXXD2

DOCUMENT TYPE: Patent
LANGUAGE: English
FAMILY ACC. NUM. COUNT: 10
PATENT INFORMATION:

| PATENT NO. | KIND | DATE | APPLICATION NO. | DATE |
|---|---|---|---|---|
| WO 9818498 | A2 | 19980507 | WO 1997-GB2958 | 19971028 <-- |
| W: AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN, CU, CZ, DE, DK, EE, ES, FI, GB, GE, GH, HU, ID, IL, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MD, MG, MK, MN, MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, UA, UG, US, UZ, VN, YU, ZW | | | | |
| RW: GH, KE, LS, MW, SD, SZ, UG, ZW, AT, BE, CH, DE, DK, ES, FI, FR, GB, GR, IE, IT | | | | |
| EP 963209 | A2 | 19991215 | EP 1997-910518 | 19971028 <-- |
| RW: DE, ES, FR, GB, IT | | | | |
| HU 9904595 | A2 | 20000428 | HU 1999-4595 | 19971028 <-- |
| US 6331289 | B1 | 20011218 | US 1997-959206 | 19971028 <-- |
| EP 1442751 | A1 | 20040804 | EP 2004-7226 | 19980424 <-- |
| R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, FI, CY | | | | |
| ES 2224379 | T3 | 20050301 | ES 1998-917461 | 19980424 <-- |
| US 2002102217 | A1 | 20020801 | US 2001-925715 | 20010810 <-- |
| US 6680047 | B2 | 20040120 | | |
| US 2005002865 | A1 | 20050106 | US 2003-734730 | 20031215 <-- |

| PRIORITY APPLN. INFO.: | | |
|---|---|---|
| GB 1996-22364 | A | 19961028 <-- |
| GB 1996-22366 | A | 19961028 <-- |
| GB 1996-22367 | A | 19961028 <-- |
| GB 1997-698 | A | 19970115 <-- |
| GB 1997-699 | A | 19970115 <-- |
| GB 1997-8265 | A | 19970424 <-- |
| GB 1997-11842 | A | 19970606 <-- |
| GB 1997-11844 | A | 19970606 <-- |
| US 1997-49264P | P | 19970606 <-- |
| GB 1996-22369 | A | 19961028 <-- |
| GB 1997-2195 | A | 19970204 <-- |
| GB 1997-11837 | A | 19970606 <-- |
| GB 1997-11839 | A | 19970606 <-- |
| US 1997-49263P | P | 19970607 <-- |
| US 1997-49266P | P | 19970607 <-- |
| US 1997-959206 | A | 19971028 <-- |
| WO 1997-GB2958 | W | 19971028 <-- |
| EP 1998-917461 | A3 | 19980424 <-- |
| US 2001-925715 | A1 | 20010810 <-- |

AB Targetable diagnostic and/or therapeutically active agents, e.g. ultrasound contrast agents, comprising a suspension in an aqueous carrier liquid of a reporter comprising gas-containing or gas-generated material, in which the reporter is coupled or linked to one or more non-bioactive vectors. Thus, a mixture of phosphatidylserine, phosphatidylcholine, and biotinamidocaproate-PEG3400-L-Ala-cholesterol (preparation given) was dispersed in 5% propylene glycol-water, flushed with perfluorobutane, and sonicated to give gas-filled encapsulated microbubbles.

L36 ANSWER 19 OF 25 HCAPLUS COPYRIGHT 2007 ACS on STN DUPLICATE 19
ACCESSION NUMBER: 1998:300863 HCAPLUS Full-text
DOCUMENT NUMBER: 129:4869
TITLE: Preparation of endothelin receptor-binding ultrasound contrast agents

INVENTOR(S): Klaveness, Jo; Naevestad, Anne; ***Cuthbertson, Alan; Solbakken, Magne***
PATENT ASSIGNEE(S): Nycomed Imaging AS, Norway; Cockbain, Julian
SOURCE: PCT Int. Appl., 98 pp.
CODEN: PIXXD2

DOCUMENT TYPE: Patent
LANGUAGE: English
FAMILY ACC. NUM. COUNT: 10
PATENT INFORMATION:

| PATENT NO. | KIND | DATE | APPLICATION NO. | DATE |
|---|---|---|---|---|
| WO 9818497 | A2 | 19980507 | WO 1997-GB2957 | 19971028 <-- |
| WO 9818497 | A3 | 19980716 | | |
| W: AL, AM, AT, AU, AZ, BB, BG, BR, BY, CA, CH, CN, CU, CZ, DE, DK, EE, ES, FI, GB, GE, GH, HU, ID, IL, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MD, MG, MN, MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, UA, UG, US, UZ, VN, YU, ZW | | | | |
| RW: GH, KE, LS, MW, SD, SZ, UG, ZW, AT, BE, CH, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG, CI, CM, GA, GN, ML, MR, NE, SN, TD, TG | | | | |
| AU 9747869 | A | 19980522 | AU 1997-47869 | 19971028 <-- |
| EP 946202 | A2 | 19991006 | EP 1997-910517 | 19971028 <-- |
| EP 946202 | B1 | 20030910 | | |
| R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, FI | | | | |
| US 6331289 | B1 | 20011218 | US 1997-959206 | 19971028 <-- |
| AT 249247 | T | 20030915 | AT 1997-910517 | 19971028 <-- |
| ES 2206689 | T3 | 20040516 | ES 1997-910517 | 19971028 <-- |
| US 2002102217 | A1 | 20020801 | US 2001-925715 | 20010810 <-- |
| US 6680047 | B2 | 20040120 | | |
| US 2005002865 | A1 | 20050106 | US 2003-734730 | 20031215 <-- |

PRIORITY APPLN. INFO.:

| | | |
|---|---|---|
| GB 1996-22364 | A | 19961028 <-- |
| GB 1996-22365 | A | 19961028 <-- |
| GB 1996-22366 | A | 19961028 <-- |
| GB 1996-22367 | A | 19961028 <-- |
| GB 1996-22368 | A | 19961028 <-- |
| GB 1996-22369 | A | 19961028 <-- |
| GB 1997-699 | A | 19970115 <-- |
| GB 1997-2195 | A | 19970204 <-- |
| GB 1997-9088 | A | 19970502 <-- |
| US 1997-48054P | P | 19970530 <-- |
| GB 1997-8265 | A | 19970424 <-- |
| GB 1997-11837 | A | 19970606 <-- |
| GB 1997-11839 | A | 19970606 <-- |
| US 1997-49264P | P | 19970606 <-- |
| US 1997-49263P | P | 19970607 <-- |
| US 1997-49266P | P | 19970607 <-- |
| US 1997-959206 | A | 19971028 <-- |
| WO 1997-GB2957 | W | 19971028 <-- |
| US 2001-925715 | A1 | 20010810 <-- |

OTHER SOURCE(S): MARPAT 129:4869

AB Compns. of matter V-L-R (V is a non-peptidic organic group having binding affinity for an endothelin receptor site; L is a linker moiety or a bond; R is a moiety detectable in in vivo ***imaging*** of a human or animal body) are described. Thus, syntheses of Gd(III) and Tc chelates of a DPTA conjugate of a lysine conjugate of 27-O-3-[2-(3-carboxyacryloylamino)-5-hydroxyphenyl]acryloyloxymyricerone are described.

L36 ANSWER 20 OF 25 HCAPLUS COPYRIGHT 2007 ACS on STN DUPLICATE 20
ACCESSION NUMBER: 1998:304262 HCAPLUS Full-text
DOCUMENT NUMBER: 129:2225
TITLE: Contrast agents
INVENTOR(S): Klaveness, Jo; Naevestad, Anne; ***Cuthbertson, Alan***
PATENT ASSIGNEE(S): Nycomed Imaging A/S, Norway; Cockbain, Julian
SOURCE: PCT Int. Appl., 99 pp.
CODEN: PIXXD2
DOCUMENT TYPE: Patent
LANGUAGE: English
FAMILY ACC. NUM. COUNT: 10
PATENT INFORMATION:

| PATENT NO. | KIND | DATE | APPLICATION NO. | DATE |
|---|---|---|---|---|
| WO 9818496 | A2 | 19980507 | WO 1997-GB2956 | 19971028 <-- |
| W: AL, AM, AT, AU, AZ, BB, BG, BR, BY, CA, CH, CN, CU, CZ, DE, DK, EE, ES, FI, GB, GE, GH, HU, ID, IL, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MD, MG, MN, MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, UA, UG, US, UZ, VN, YU, ZW | | | | |
| RW: GH, KE, LS, MW, SD, SZ, UG, ZW, AT, BE, CH, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG, CI, CM, GA, GN, ML, MR, NE, SN, TD, TG | | | | |
| AU 9747868 | A | 19980522 | AU 1997-47868 | 19971028 <-- |
| EP 971747 | A2 | 20000119 | EP 1997-910516 | 19971028 <-- |
| EP 971747 | B1 | 20051228 | | |
| R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, FI | | | | |
| US 6331289 | B1 | 20011218 | US 1997-959206 | 19971028 <-- |
| AT 314097 | T | 20060115 | AT 1997-910516 | 19971028 <-- |
| ES 2257771 | T3 | 20060801 | ES 1997-910516 | 19971028 <-- |
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| US 2001016587 | A1 | 20010823 | US 2001-785177 | 20010220 <-- |
| US 6524552 | B2 | 20030225 | | |
| US 2002102217 | A1 | 20020801 | US 2001-925715 | 20010810 <-- |
| US 6680047 | B2 | 20040120 | | |
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| US 6921525 | B2 | 20050726 | | |
| US 2005002865 | A1 | 20050106 | US 2003-734730 | 20031215 <-- |
| US 2005201930 | A1 | 20050915 | US 2005-108598 | 20050418 <-- |
| US 7182934 | B2 | 20070227 | | |

| PRIORITY APPLN. INFO.: | | | |
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| | GB 1996-22364 | A | 19961028 <-- |
| | GB 1996-22365 | A | 19961028 <-- |
| | GB 1996-22366 | A | 19961028 <-- |
| | GB 1996-22367 | A | 19961028 <-- |
| | GB 1996-22368 | A | 19961028 <-- |
| | GB 1996-22369 | A | 19961028 <-- |
| | GB 1997-699 | A | 19970115 <-- |
| | GB 1997-2195 | A | 19970204 <-- |
| | GB 1997-6063 | A | 19970324 <-- |
| | US 1997-58247P | P | 19970909 <-- |
| | GB 1997-8265 | A | 19970424 <-- |
| | GB 1997-11837 | A | 19970606 <-- |
| | GB 1997-11839 | A | 19970606 <-- |
| | US 1997-49264P | P | 19970606 <-- |
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| | US 1997-959206 | A | 19971028 <-- |
| | WO 1997-GB2956 | W | 19971028 <-- |
| | US 1999-300434 | A3 | 19990428 <-- |
| | US 2001-785177 | A3 | 20010220 <-- |
| | US 2001-925715 | A1 | 20010810 <-- |
| | US 2003-370092 | A3 | 20030221 <-- |

OTHER SOURCE(S): MARPAT 129:2225

AB The invention provides a composition of matter (I): V-L-R where V is an organic group having binding affinity for an angiotensin II receptor site, L is a linker moiety or a bond, and R is a moiety detectable in in vivo ***imaging*** of a human or animal body, with the provisos that where V is angiotensin or a peptidic angiotensin derivative or analog then V-L-R is other than a nonmetal radionuclide substituted peptide (e.g. 125I substituted angiotensin II) and L-V is other than simply a peptide with a chelating agent amide bonded to a side chain thereof. This composition of matter may be used to image cardiovascular diseases and disorders.

L36 ANSWER 21 OF 25 HCAPLUS COPYRIGHT 2007 ACS on STN

ACCESSION NUMBER: 2007:173633 HCAPLUS Full-text

TITLE: Separation processes

INVENTOR(S): Rongved, Pal; Loevhaug, Dagfinn; Fjerdingstad, Hege; ***Solbakken, Magne***; Godal, Aslak; ***Cuthbertson, Alan***

PATENT ASSIGNEE(S): Norway

SOURCE: U.S. Pat. Appl. Publ., 17pp., Cont.-in-part of U.S. Ser. No. 722,075.
CODEN: USXXCO

DOCUMENT TYPE: Patent

LANGUAGE: English

FAMILY ACC. NUM. COUNT: 10

PATENT INFORMATION:

| PATENT NO. | KIND | DATE | APPLICATION NO. | DATE |
|---|---|---|---|---|
| US 2007036722 | A1 | 20070215 | US 2006-498651 | 20060803 <-- |
| CN 1234742 | A | 19991110 | CN 1997-199047 | 19971028 <-- |
| HU 9904595 | A2 | 20000428 | HU 1999-4595 | 19971028 <-- |
| US 6331289 | B1 | 20011218 | US 1997-959206 | 19971028 <-- |
| AT 318618 | T | 20060315 | AT 1997-910514 | 19971028 <-- |
| US 6261537 | B1 | 20010717 | US 1997-960054 | 19971029 <-- |
| EP 1442751 | A1 | 20040804 | EP 2004-7226 | 19980424 <-- |
| R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, FI, CY | | | | |
| ES 2224379 | T3 | 20050301 | ES 1998-917461 | 19980424 <-- |
| KR 2000052829 | A | 20000825 | KR 1999-703658 | 19990427 <-- |
| US 2002102215 | A1 | 20020801 | US 2001-765614 | 20010122 <-- |
| US 2002102217 | A1 | 20020801 | US 2001-925715 | 20010810 <-- |
| US 6680047 | B2 | 20040120 | | |
| CN 1440816 | A | 20030910 | CN 2002-160420 | 20021230 <-- |
| US 2004141922 | A1 | 20040722 | US 2003-722075 | 20031126 <-- |
| US 2005002865 | A1 | 20050106 | US 2003-734730 | 20031215 <-- |

PRIORITY APPLN. INFO.:

| | | |
|---|---|---|
| GB 1996-22366 | A | 19961028 <-- |
| GB 1996-22367 | A | 19961028 <-- |
| GB 1996-22368 | A | 19961028 <-- |
| GB 1997-699 | A | 19970115 <-- |
| GB 1997-8265 | A | 19970424 <-- |
| GB 1997-11842 | A | 19970606 <-- |
| GB 1997-11846 | A | 19970606 <-- |
| US 1997-49264P | P | 19970606 <-- |

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| US 1997-49265P | P | 19970606 <-- |
| US 1997-49268P | P | 19970606 <-- |
| US 1997-958993 | A2 | 19971028 <-- |
| US 1997-960054 | A1 | 19971029 <-- |
| US 2001-765614 | B1 | 20010122 <-- |
| US 2003-722075 | A2 | 20031126 <-- |
| GB 1996-22369 | A | 19961028 <-- |
| GB 1997-2195 | A | 19970204 <-- |
| GB 1997-11837 | A | 19970606 <-- |
| GB 1997-11839 | A | 19970606 <-- |
| US 1997-49263P | P | 19970607 <-- |
| US 1997-49266P | P | 19970607 <-- |
| US 1997-959206 | A | 19971028 <-- |
| EP 1998-917461 | A3 | 19980424 <-- |
| US 2001-925715 | A1 | 20010810 <-- |

AB Separation of target material from a liquid sample is achieved by coupling the target to targetable encapsulated gas microbubbles, allowing the microbubbles and coupled target to float to the surface of the sample to form a floating microbubble/target layer, and separating this layer from the sample. In a pos. separation process the microbubbles are then removed from the target, e.g. by bursting. In a neg. separation process target-free sample material is recovered following separation of the floating layer. The method may also be used diagnostically to detect the presence of a disease marker in a sample. Novel separation apparatus is also described. Gas microbubbles encapsulated with DSPS and thiolated anti-CD34 antibodies-Mal-PEG2000DSPE, useful for separation of hematopoietic stem cells, were prepared

L36 ANSWER 22 OF 25 HCAPLUS COPYRIGHT 2007 ACS on STN
ACCESSION NUMBER: 2004:780565 HCAPLUS Full-text
DOCUMENT NUMBER: 141:277892
TITLE: Methods of radiofluorination of peptides and other biologically active vectors
INVENTOR(S): ***Cuthbertson, Alan; Solbakken, Magne*** ; Arukwe, Joseph Maduabuchi; Karlsen, Hege; Glaser, Matthias Eberhard
PATENT ASSIGNEE(S): Amersham Health AS, Norway; Hammersmith Imanet Ltd.
SOURCE: PCT Int. Appl., 56 pp.
CODEN: PIXXD2
DOCUMENT TYPE: Patent
LANGUAGE: English
FAMILY ACC. NUM. COUNT: 1
PATENT INFORMATION:

| PATENT NO. | KIND | DATE | APPLICATION NO. | DATE |
|---|---|---|---|---|
| WO 2004080492 | A1 | 20040923 | WO 2004-GB1052 | 20040312 <-- |
| W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BW, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, EG, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NA, NI, NO, NZ, OM, PG, PH, PL, PT, RO, RU, SC, SD, SE, SG, SK, SL, SY, TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, YU, ZA, ZM, ZW | | | | |
| RW: BW, GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM, AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE, IT, LU, MC, NL, PL, PT, RO, SE, SI, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG | | | | |
| AU 2004218879 | A1 | 20040923 | AU 2004-218879 | 20040312 <-- |
| CA 2518889 | A1 | 20040923 | CA 2004-2518889 | 20040312 <-- |

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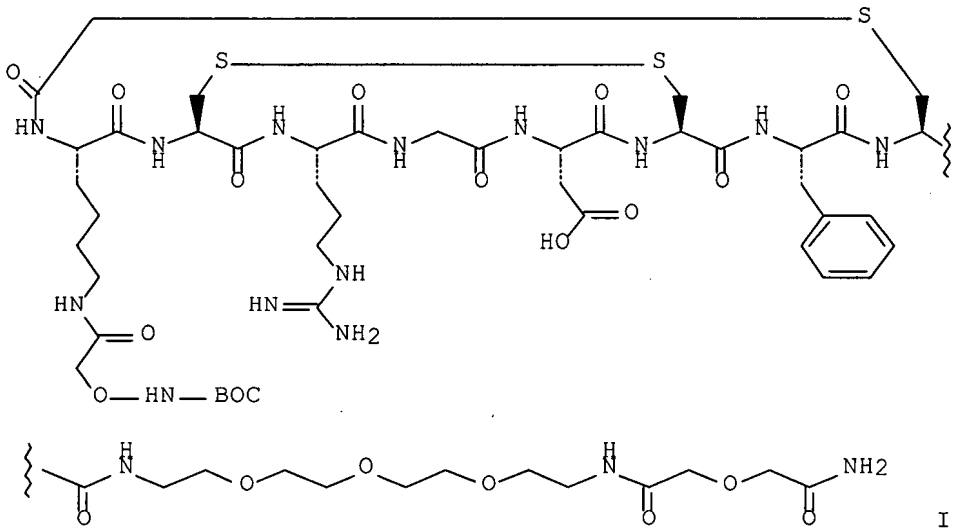
| EP 1601384 | A1 | 20051207 | EP 2004-720084 | | 20040312 <-- |
| R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, LT, LV, FI, RO, MK, CY, AL, TR, BG, CZ, EE, HU, PL, SK | | | | | |
| BR 2004008236 | A | 20060301 | BR 2004-8236 | | 20040312 <-- |
| CN 1758925 | A | 20060412 | CN 2004-80006725 | | 20040312 <-- |
| JP 2006523658 | T | 20061019 | JP 2006-505951 | | 20040312 <-- |
| NO 2005004185 | A | 20051110 | NO 2005-4185 | | 20050909 <-- |
| PRIORITY APPLN. INFO.: | | | GB 2003-5704 | A | 20030313 <-- |
| | | | WO 2004-GB1052 | A | 20040312 |

OTHER SOURCE(S): MARPAT 141:277892
GI



I

AB The invention relates to diagnostic and radio-diagnostic agents, including biol. active vectors labeled with positron-emitting nuclides. It further relates to methods and reagents for (18F)-fluorination of vectors, where a vector is defined as a mol. with an affinity for a specific biol. target, and is preferably a peptide. The resultant 18F-labeled conjugates are useful as radiopharmaceuticals, specifically for use in Positron Emission Tomog. (PET). Compds. 18F-linker-X-N:C(Y)-vector and vector-X-N:C(Y)- linker-18F (X is CONH, NH, O, NHCONH or NHCSNH; Y is H, alkyl or aryl) are claimed. Thus, I (Boc = tert-butoxycarbonyl), prepared by standard peptide synthesis and coupling with Boc-NHOCH2CO2H, was deprotected and conjugated with 4-18FC6H4CHO.

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| Referenced Author (RAU) | Year (RPY) | VOL (RVL) | PG (RPG) | Referenced Work (RWK) | Referenced File |
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L36 ANSWER 23 OF 25 HCAPLUS COPYRIGHT 2007 ACS on STN
ACCESSION NUMBER: 2003:777733 HCAPLUS Full-text
DOCUMENT NUMBER: 139:277170
TITLE: Methods and reagents for radiofluorination, particularly of peptides

INVENTOR(S): ***Cuthbertson, Alan; Solbakken, Magne*** ; Arukwe, Joseph Maduabuchi; Karlsen, Hege
PATENT ASSIGNEE(S): Amersham PLC, UK
SOURCE: PCT Int. Appl., 38 pp.
CODEN: PIXXD2
DOCUMENT TYPE: Patent
LANGUAGE: English
FAMILY ACC. NUM. COUNT: 1
PATENT INFORMATION:

| PATENT NO. | KIND | DATE | APPLICATION NO. | DATE |
|---|---|---|---|---|
| WO 2003080544 | A1 | 20031002 | WO 2003-GB1332 | 20030320 <-- |
| W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NI, NO, NZ, OM, PH, PL, PT, RO, RU, SC, SD, SE, SG, SK, SL, TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, YU, ZA, ZM, ZW | | | | |
| RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM, AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE, IT, LU, MC, NL, PT, RO, SE, SI, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG | | | | |
| AU 2003214446 | A1 | 20031008 | AU 2003-214446 | 20030320 <-- |
| EP 1487762 | A1 | 20041222 | EP 2003-710021 | 20030320 <-- |
| EP 1487762 | B1 | 20070228 | | |
| R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, LT, LV, FI, RO, MK, CY, AL, TR, BG, CZ, EE, HU, SK | | | | |
| US 2005142061 | A1 | 20050630 | US 2003-508682 | 20030320 <-- |
| JP 2005520857 | T | 20050714 | JP 2003-578305 | 20030320 <-- |
| PRIORITY APPLN. INFO.: | | | GB 2002-6750 A | 20020322 <-- |
| | | | WO 2003-GB1332 W | 20030320 <-- |

OTHER SOURCE(S): MARPAT 139:277170
GI

* STRUCTURE DIAGRAM TOO LARGE FOR DISPLAY - AVAILABLE VIA OFFLINE PRINT *

AB The invention relates to the synthesis of 18F-labeled compds., particularly peptides, for use as radiopharmaceuticals, specifically for use in positron emission tomog. (PET). The radiofluorination method involves reacting a compound X-CH2CONH-peptide (X is a halogen leaving group, preferably chloro) or maleimido-Y-CONH-peptide (Y is a C1-10 hydrocarbyl group optionally containing 1-6 heteroatoms) with a compound 18F-(linker)-SH, in which the linker is a C1-30 hydrocarbyl group optionally containing 1-10 heteroatoms. Thus, compound I was prepared by site-specific conjugation of 4-FCH2C6H4CONHCH2CH2SCPh3 to the maleimide-modified peptide.

RETABLE

| Referenced Author (RAU) | Year (RPY) | VOL (RVL) | PG (RPG) | Referenced Work (RWK) | Referenced File |
|---|---|---|---|---|---|
| Dean, R | 1992 | | | US 5144043 A | HCAPLUS |
| Griffiths, G | 1999 | | | WO 9911590 A | HCAPLUS |
| Hwang, D | 1991 | 32 | 1730 | JOURNAL OF NUCLEAR M | HCAPLUS |

L36 ANSWER 24 OF 25 HCAPLUS COPYRIGHT 2007 ACS on STN
ACCESSION NUMBER: 1999:708880 HCAPLUS Full-text
DOCUMENT NUMBER: 131:319884

TITLE: Targetable encapsulated gas microbubbles for separation of target material from liquid samples and separation apparatus

INVENTOR(S): ***Cuthbertson, Alan***; Rongved, Pal; Lovhaug, Dagfinn; Fjerdingstad, Hege; ***Solbakken, Magne***; Godal, Aslak

PATENT ASSIGNEE(S): Nycomed Imaging As, Norway

SOURCE: PCT Int. Appl., 54 pp.
CODEN: PIXXD2

DOCUMENT TYPE: Patent

LANGUAGE: English

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

| PATENT NO. | KIND | DATE | APPLICATION NO. | DATE |
|---|---|---|---|---|
| WO 9955837 | A2 | 19991104 | WO 1999-GB1317 | 19990428 <-- |
| WO 9955837 | A3 | 20000210 | | |
| W: AE, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN, CU, CZ, DE, DK, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MD, MG, MK, MN, MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, UA, UG, US, UZ, VN, YU, ZA, ZW | | | | |
| RW: GH, GM, KE, LS, MW, SD, SL, SZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG | | | | |
| CA 2326386 | A1 | 19991104 | CA 1999-2326386 | 19990428 <-- |
| AU 9937197 | A | 19991116 | AU 1999-37197 | 19990428 <-- |
| EP 1073716 | A2 | 20010207 | EP 1999-919396 | 19990428 <-- |
| EP 1073716 | B1 | 20040428 | | |
| R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, FI | | | | |
| JP 2002512886 | T | 20020508 | JP 2000-545981 | 19990428 <-- |
| AT 265525 | T | 20040515 | AT 1999-919396 | 19990428 <-- |
| IN 2000MN00515 | A | 20050715 | IN 2000-MN515 | 20001018 <-- |
| NO 2000005383 | A | 20001213 | NO 2000-5383 | 20001026 <-- |
| US 2003104359 | A1 | 20030605 | US 2002-294598 | 20021115 <-- |

PRIORITY APPLN. INFO.:

| | | |
|---|---|---|
| GB 1998-9083 | A | 19980428 <-- |
| GB 1998-9085 | A | 19980428 <-- |
| US 1998-85819P | P | 19980518 <-- |
| US 1998-85826P | P | 19980518 <-- |
| WO 1999-GB1317 | W | 19990428 <-- |
| US 2000-694893 | B1 | 20001025 <-- |

AB Separation of target material from a liquid sample is achieved by coupling the target to targetable encapsulated gas microbubbles, allowing the microbubbles and coupled target to float to the surface of the sample to form a floating microbubble/target layer, and separating this layer from the sample. In a pos. separation process the microbubbles are then removed from the target, e.g. by bursting. In a neg. separation process target-free sample material is recovered following separation of the floating layer. The method may also be used diagnostically to detect the presence of a disease marker in a sample. Novel separation apparatus is also described. Perfluorobutane gas microbubbles encapsulated with distearoylphosphatidylserine doped with Mal-PEG2000-distearoylphosphatidylethanolamine (DSPE) was prepared and reacted with thiolated anti-CD34 antibodies to make a reagent useful for separating CD34-pos. cells.

L36 ANSWER 25 OF 25 WPIX COPYRIGHT 2007 THE THOMSON CORP on STN
ACCESSION NUMBER: 2004-625405 [60] WPIX

DOC. NO. CPI: C2004-224919 [60]
TITLE: Diagnostic ***imaging agent***, useful for diagnostic imaging of cardiovascular and inflammatory diseases, comprises a matrix metalloproteinase inhibitor labeled with a gamma-emitting radionuclide
DERWENT CLASS: B04; B05; K08
INVENTOR: ARUKWE J; BREYHOLZ H; ***CUTHBERTSON A***; DAVIS J; HEYWOOD K; KOPKA K; LEVKAU B; MENDIZABAL M; RICKETTS S; SCHAFERS M; STOREY A; STOREY A E; WAGNER S; WILSON I; WYNN D
PATENT ASSIGNEE: (AMSH-C) AMERSHAM PLC; (GENE-C) GE HEALTHCARE LTD
COUNTRY COUNT: 107

PATENT INFO ABBR.:

| PATENT NO | KIND | DATE | WEEK | LA | PG | MAIN IPC |
|---|---|---|---|---|---|---|
| WO 2004069365 | A1 | 20040819 | (200460)* | EN | 61[12] | |
| AU 2004210208 | A1 | 20040819 | (200559) | EN | | |
| EP 1592458 | A1 | 20051109 | (200573) | EN | | |
| NO 2005003776 | A | 20050930 | (200574) | NO | | |
| CN 1747749 | A | 20060315 | (200649) | ZH | | |
| JP 2006519216 | W | 20060824 | (200656) | JA | 57 | |

APPLICATION DETAILS:

| PATENT NO | KIND | APPLICATION | DATE |
|---|---|---|---|
| WO 2004069365 | A1 | WO 2004-GB524 | 20040210 |
| AU 2004210208 | A1 | AU 2004-210208 | 20040210 |
| CN 1747749 | A | CN 2004-80003837 | 20040210 |
| EP 1592458 | A1 | EP 2004-709657 | 20040210 |
| EP 1592458 | A1 | WO 2004-GB524 | 20040210 |
| NO 2005003776 | A | NO 2005-3776 | 20050809 |
| JP 2006519216 | W | WO 2004-GB524 | 20040210 |
| JP 2006519216 | W | JP 2006-502261 | 20040210 |

FILING DETAILS:

| PATENT NO | KIND | | PATENT NO | |
|---|---|---|---|---|
| AU 2004210208 | A1 | Based on | WO 2004069365 | A |
| EP 1592458 | A1 | Based on | WO 2004069365 | A |
| JP 2006519216 | W | Based on | WO 2004069365 | A |

PRIORITY APPLN. INFO: ***GB 2003-7524 20030401***
***GB 2003-2891 20030210***
AN 2004-625405 [60] WPIX
AB WO 2004069365 A1 UPAB: 20060203
NOVELTY - Diagnostic ***imaging agent*** (A) comprises a matrix metalloproteinase inhibitor (I) labeled with a gamma-emitting radionuclide.
DETAILED DESCRIPTION - Diagnostic ***imaging agent*** (A) comprises a matrix metalloproteinase inhibitor of formula (I) labeled with a gamma-emitting radionuclide.
Either R1 = H, OH, 1-6C alkyl, 6-14C aryl or 7-20C arylalkyl; or
C+R1+R5 = 6-8C cycloalkyl ring or a 4-6C heterocyclic ring; or
C+R1+R4 = 4-6C heterocyclic ring containing 5-7 atoms and 1 or 2 heteroatoms from N or O;
R2, R3 = H, OH, halo, 1-6C alkyl, 1-6C alkoxy, 1-6C amino, 6-14C aryl, 7-20C arylalkyl or 7-20C carbamoylaryl;

R4 = 6-14C aryl, 4-6C heteroaryl, 7-20C arylalkyl, 7-20C carbamoylaryl or arylcarbamoylaryl; and

R5 = H or 1-6C alkyl.

INDEPENDENT CLAIMS are also included for:

(1) a ligand conjugate which comprises (I), conjugated to a ligand suitable for the co-ordination of a gamma-emitting radio metal (99m-Tc, 111ln, 113mln, 67Cu or 67Ga);

(2) a precursor useful in the preparation of (A) comprising a group suitable for reaction with a gamma-emitting isotope of iodine to give (A);

(3) a pharmaceutical composition comprising (A) with a biocompatible carrier; and

(4) a kit for the preparation of a pharmaceutical composition of (A).

ACTIVITY - Cardiovascular-Gen.; Antiarteriosclerotic; Antiinflammatory; Respiratory-Gen.

MECHANISM OF ACTION - Matrix metalloproteinase (MMP) inhibitor.

Compounds (I) were tested for their MMP inhibitory activity using biological assays. The result showed that the median inhibitory concentration value of (2R)-N-hydroxy-2-(((4-iodophenyl)sulfonyl)(pyridin- 3-ylmethyl)amino)-3-methylbutanamide was 2.5 nM.

USE - (A) is useful for the diagnostic imagining of cardiovascular disease (preferably atherosclerosis and congestive heart failure) and inflammatory diseases (preferably chronic obstructive pulmonary disease) (claimed).

=> d que l9
L1 STR

* STRUCTURE DIAGRAM TOO LARGE FOR DISPLAY - AVAILABLE VIA OFFLINE PRINT *

Structure attributes must be viewed using STN Express query preparation.
L3 1 SEA FILE=HCAPLUS ABB=ON PLU=ON US2006-566487/AP
L5 4 SEA FILE=REGISTRY SSS FUL L1
L8 1 SEA FILE=HCAPLUS ABB=ON PLU=ON L5
L9 1 SEA FILE=HCAPLUS ABB=ON PLU=ON (L8 OR L3)

=> d ibib abs hitstr retable l9 tot

L9 ANSWER 1 OF 1 HCAPLUS COPYRIGHT 2007 ACS on STN
ACCESSION NUMBER: 2005:120957 HCAPLUS Full-text
DOCUMENT NUMBER: 142:219561
TITLE: Preparation of peptide-based compounds as diagnostic imaging agents
INVENTOR(S): Cuthbertson, Alan; Solbakken, Magne
PATENT ASSIGNEE(S): Amersham Health AS, Norway
SOURCE: PCT Int. Appl., 40 pp.
CODEN: PIXXD2
DOCUMENT TYPE: Patent
LANGUAGE: English
FAMILY ACC. NUM. COUNT: 1
PATENT INFORMATION:

instant application

| PATENT NO. | KIND | DATE | APPLICATION NO. | DATE |
|---|---|---|---|---|
| WO 2005012335 | A1 | 20050210 | WO 2004-GB3150 | 20040721 |

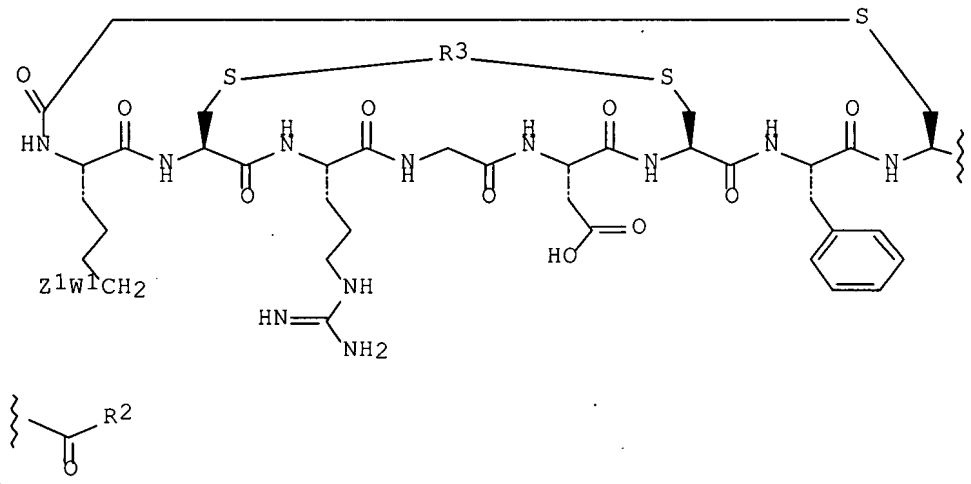
W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BW, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, EG, ES, FI, GB, GD,

GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC,
LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NA, NI,
NO, NZ, OM, PG, PH, PL, PT, RO, RU, SC, SD, SE, SG, SK, SL, SY,
TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, YU, ZA, ZM, ZW
RW: BW, GH, GM, KE, LS, MW, MZ, NA, SD, SL, SZ, TZ, UG, ZM, ZW, AM,
AZ, BY, KG, KZ, MD, RU, TJ, TM, AT, BE, BG, CH, CY, CZ, DE, DK,
EE, ES, FI, FR, GB, GR, HU, IE, IT, LU, MC, NL, PL, PT, RO, SE,
SI, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE,
SN, TD, TG

| | | | | | |
|---|---|---|---|---|---|
| CA 2533321 | A1 | 20050210 | CA 2004-2533321 | | 20040721 |
| EP 1648925 | A1 | 20060426 | EP 2004-743485 | | 20040721 |
| R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, FI, RO, CY, TR, BG, CZ, EE, HU, PL, SK | | | | | |
| CN 1829735 | A | 20060906 | CN 2004-80022044 | | 20040721 |
| BR 2004012986 | A | 20061003 | BR 2004-12986 | | 20040721 |
| HU 200600230 | A2 | 20070129 | HU 2006-230 | | 20040721 |
| US 2006193773 | A1 | 20060831 | US 2006-566487 | | 20060130 <-- |
| NO 2006000825 | A | 20060329 | NO 2006-825 | | 20060221 |
| PRIORITY APPLN. INFO.: | | | GB 2003-17815 | A | 20030730 |
| | | | WO 2004-GB3150 | W | 20040721 |

OTHER SOURCE(S): CASREACT 142:219561; MARPAT 142:219561

GI




I

AB The invention relates to compds. I [R2 is [NH(CH2CH2O)3CH2CH2NHCOCH2OCH2CO ]0-10NH2; R3 is an alkylene or alkenylene bridge; W1 is absent or a spacer moiety (hetero)hydrocarbyl preferably derived from glutaric and/or succinic acid and/or a polyethylene glycol-based unit and/or a unit [NH(CH2CH2O)3CH2CH2NHCOCH2OCH2CO]n; Z1 is an antineoplastic agent, a chelating agent or a reporter moiety] and their use as targeting vectors that bind to receptors associated with angiogenesis. Compds. I may thus be used for diagnosis or therapy of various diseases. Thus, compound I [R2 is NH2, R3 is CH2, Z1-W1 is FCH2CH2SCH2CONH(CH2CH2O)5CH2CH2NHCOCH2OCH2CONH] was prepared via the solid-phase method and showed Ki = 7 nmol in an αvβ3 integrin receptor binding assay.

IT *840474-71-7P 840474-72-8P*
RL: DGN (Diagnostic use); SPN (Synthetic preparation); BIOL (Biological study); PREP (Preparation); USES (Uses)
(preparation of peptide-based compds. as diagnostic imaging agents)
RN 840474-71-7 HCAPLUS
CN L-Cysteinamide, N6-(30-fluoro-1,5,25-trioxo-3,9,12,15,18,21-hexaoxa-27-thia-6,24-diazatriacont-1-yl)-N2-(mercaptoacetyl)-L-lysyl-S-(mercaptomethyl)-L-cysteinyl-L-arginylglycyl-L-α-aspartyl-L-cysteinyl-L-phenylalanyl-, cyclic (1→8),(2→6)-bis(thioether) (9CI) (CA INDEX NAME)

*** STRUCTURE DIAGRAM IS NOT AVAILABLE ***
RN 840474-72-8 HCAPLUS
CN L-Cysteinamide, N6-[30-(fluoro-18F)-1,5,25-trioxo-3,9,12,15,18,21-hexaoxa-27-thia-6,24-diazatriacont-1-yl]-N2-(mercaptoacetyl)-L-lysyl-S-(mercaptomethyl)-L-cysteinyl-L-arginylglycyl-L-α-aspartyl-L-cysteinyl-L-phenylalanyl-, cyclic (1→8),(2→6)-bis(thioether) (9CI) (CA INDEX NAME)

*** STRUCTURE DIAGRAM IS NOT AVAILABLE ***
IT *840474-69-3P 840474-70-6P*
RL: RCT (Reactant); SPN (Synthetic preparation); PREP (Preparation); RACT (Reactant or reagent)
(preparation of peptide-based compds. as diagnostic imaging agents)
RN 840474-69-3 HCAPLUS
CN L-Cysteinamide, N2-(mercaptoacetyl)-L-lysyl-S-(mercaptomethyl)-L-cysteinyl-L-arginylglycyl-L-α-aspartyl-L-cysteinyl-L-phenylalanyl-, cyclic (1→8),(2→6)-bis(thioether) (9CI) (CA INDEX NAME)

*** STRUCTURE DIAGRAM IS NOT AVAILABLE ***
RN 840474-70-6 HCAPLUS
CN L-Cysteinamide, N6-(26-chloro-1,5,25-trioxo-3,9,12,15,18,21-hexaoxa-6,24-diazahexacos-1-yl)-N2-(mercaptoacetyl)-L-lysyl-S-(mercaptomethyl)-L-cysteinyl-L-arginylglycyl-L-α-aspartyl-L-cysteinyl-L-phenylalanyl-, cyclic (1→8),(2→6)-bis(thioether) (9CI) (CA INDEX NAME)

*** STRUCTURE DIAGRAM IS NOT AVAILABLE ***
RETABLE

| Referenced Author (RAU) | Year (RPY) | VOL (RVL) | PG (RPG) | Referenced Work (RWK) | Referenced File |
|---|---|---|---|---|---|
| Bonasera, T | 2002 | | | WO 02062819 A | HCAPLUS |
| Harris, T | 1996 | 6 | 1741 | BIOORGANIC & MEDICIN | HCAPLUS |
| Indrevoll, B | 2001 | | | WO 0177145 A | HCAPLUS |
| Indrevoll, B | 2003 | | | WO 03006491 A | HCAPLUS |
| Lister-James, J | 1999 | | | US 5888474 A | HCAPLUS |
| Pearson, D | 1996 | 39 | 1372 | JOURNAL OF MEDICINAL | HCAPLUS |
| Srinivasan, A | 2002 | | | WO 0220610 A | HCAPLUS |

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(FILE 'HOME' ENTERED AT 13:43:12 ON 15 MAR 2007)

FILE 'REGISTRY' ENTERED AT 13:43:18 ON 15 MAR 2007
L1 STRUCTURE UPLOADED
L2 0 SEA SSS SAM L1

FILE 'HCAPLUS' ENTERED AT 13:43:41 ON 15 MAR 2007

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L3            1 SEA ABB=ON  PLU=ON  US2006-566487/AP
                D SCAN
                SEL RN L3

     FILE 'REGISTRY' ENTERED AT 13:44:00 ON 15 MAR 2007
L4           11 SEA ABB=ON  PLU=ON  (100360-56-3/BI OR 19721-22-3/BI OR
                541-88-8/BI OR 606975-95-5/BI OR 606975-96-6/BI OR 606976-02-7/
                BI OR 840474-68-2/BI OR 840474-69-3/BI OR 840474-70-6/BI OR
                840474-71-7/BI OR 840474-72-8/BI)
                D SCAN
                D QUE L1
L5            4 SEA SSS FUL L1
                D SCAN
                SAVE L5 YOUNG487/A TEMP
L6            4 SEA ABB=ON  PLU=ON  L4 AND L5
                D SCAN
L7            7 SEA ABB=ON  PLU=ON  L4 NOT L6
                D SCAN

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L8            1 SEA ABB=ON  PLU=ON  L5
L9            1 SEA ABB=ON  PLU=ON  (L8 OR L3)

     FILE 'BEILSTEIN' ENTERED AT 13:46:21 ON 15 MAR 2007
L10           0 SEA SSS FUL L1

     FILE 'MARPAT' ENTERED AT 13:46:33 ON 15 MAR 2007
L11           1 SEA SSS SAM L1
L12           2 SEA SSS FUL L1
L13           1 SEA ABB=ON  PLU=ON  L12/COM
L14           0 SEA ABB=ON  PLU=ON  L13 NOT L9

     FILE 'HCAPLUS' ENTERED AT 13:48:14 ON 15 MAR 2007
                E CUTHBERTSON A/AU
L15          94 SEA ABB=ON  PLU=ON  ("CUTHBERTSON A"/AU OR "CUTHBERTSON A
                C"/AU OR "CUTHBERTSON A F"/AU OR "CUTHBERTSON A F J"/AU OR
                "CUTHBERTSON A G S"/AU OR "CUTHBERTSON A M"/AU OR "CUTHBERTSON
                A S"/AU OR "CUTHBERTSON A Z"/AU OR "CUTHBERTSON ALAN"/AU OR
                "CUTHBERTSON ALAN J S"/AU OR "CUTHBERTSON ALAN S"/AU OR
                "CUTHBERTSON ALLAN S"/AU)
                E SOLBAKKEN M/AU
L16          28 SEA ABB=ON  PLU=ON  ("SOLBAKKEN M"/AU OR "SOLBAKKEN MAGNE"/AU)

L17          19 SEA ABB=ON  PLU=ON  L15 AND L16
L18          14 SEA ABB=ON  PLU=ON  L17 AND (AY<2004 OR PY<2004 OR PRY<2004)
L19          32 SEA ABB=ON  PLU=ON  (L15 OR L16) AND (IMAGING?)
L20          20 SEA ABB=ON  PLU=ON  L19 AND (AY<2004 OR PY<2004 OR PRY<2004)
L21          24 SEA ABB=ON  PLU=ON  (L20 OR L18)

     FILE 'HCAPLUS, MEDLINE, EMBASE, BIOSIS, DRUGU, WPIX' ENTERED AT 13:50:28
     ON 15 MAR 2007
L22         437 SEA ABB=ON  PLU=ON  CUTHBERTSON A?/AU
L23          74 SEA ABB=ON  PLU=ON  SOLBAKKEN M?/AU
L24          49 SEA ABB=ON  PLU=ON  L22 AND L23
L25          36 SEA ABB=ON  PLU=ON  L24 AND IMAGING?
L26          25 SEA ABB=ON  PLU=ON  L25 AND (AY<2004 OR PY<2004 OR PRY<2004)
L27          47 SEA ABB=ON  PLU=ON  (L22 OR L23) AND (IMAGING AGENT?)
L28          29 SEA ABB=ON  PLU=ON  L27 AND (AY<2004 OR PY<2004 OR PRY<2004)
L29          40 SEA ABB=ON  PLU=ON  (L26 OR L28)
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L30          29 SEA ABB=ON  PLU=ON  L29 AND (IMAGING AGENT?)

     FILE 'STNGUIDE' ENTERED AT 13:53:33 ON 15 MAR 2007

     FILE 'REGISTRY' ENTERED AT 13:55:15 ON 15 MAR 2007
               D RSD L6 TOT
L31            4 SEA ABB=ON  PLU=ON  105465.1/RID
L32            4 SEA ABB=ON  PLU=ON  L31 AND 46.150/RID
L33            4 SEA ABB=ON  PLU=ON  (L31 OR L32)
L34            4 SEA ABB=ON  PLU=ON  (L6 OR L33)

     FILE 'MEDLINE, EMBASE, BIOSIS, DRUGU, CAOLD' ENTERED AT 13:57:13 ON 15
     MAR 2007
L35            0 SEA ABB=ON  PLU=ON  L5

     FILE 'STNGUIDE' ENTERED AT 13:57:23 ON 15 MAR 2007
               D QUE L21
               D QUE L30

     FILE 'HCAPLUS, MEDLINE, EMBASE, BIOSIS, WPIX' ENTERED AT 13:57:33 ON 15
     MAR 2007
L36           25 DUP REM L21 L30 (28 DUPLICATES REMOVED)
                    ANSWERS '1-24' FROM FILE HCAPLUS
                    ANSWER '25' FROM FILE WPIX
               D IBIB ABS HITSTR RETABLE L36 TOT
               D QUE L9
               D IBIB ABS HITSTR RETABLE L9 TOT


     FILE HOME

     FILE REGISTRY
     Property values tagged with IC are from the ZIC/VINITI data file
     provided by InfoChem.

     STRUCTURE FILE UPDATES:   14 MAR 2007  HIGHEST RN 926494-79-3
     DICTIONARY FILE UPDATES:  14 MAR 2007  HIGHEST RN 926494-79-3

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     TSCA INFORMATION NOW CURRENT THROUGH December 2, 2006

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     http://www.cas.org/ONLINE/UG/regprops.html

     FILE HCAPLUS

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     the American Chemical Society and is provided to assist you in searching
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FILE COVERS 1907 - 15 Mar 2007 VOL 146 ISS 12
FILE LAST UPDATED: 14 Mar 2007 (20070314/ED)

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This file contains CAS Registry Numbers for easy and accurate substance identification.

FILE BEILSTEIN
FILE LAST UPDATED ON JANUARY 10, 2007

FILE COVERS 1771 TO 2006.
***FILE CONTAINS 9,780,003 SUBSTANCES***

>>>PLEASE NOTE: Reaction Data and substance data are stored in separate documents and can not be searched together in one query. Reaction data for BEILSTEIN compounds may be displayed immediately with the display codes PRE (preparations) and REA (reactions). A substance answer set retrieved after the search for a chemical name, a compounds with available reaction information by combining with PRE/FA, REA/FA or more generally with RX/FA. The BEILSTEIN Registry Number (BRN) is the link between a BEILSTEIN compound and belonging reactions. For mo detailed reaction searches BRNs can be searched as reaction partner BRNs Reactant BRN (RX.RBRN) or Product BRN (RX.PBRN).<<<

>>> FOR SEARCHING PREPARATIONS SEE HELP PRE <<<

**********
* PLEASE NOTE THAT THERE ARE NO FORMATS FREE OF COST. *
* SET NOTICE FEATURE: THE COST ESTIMATES CALCULATED FOR SET NOTICE *
* ARE BASED ON THE HIGHEST PRICE CATEGORY. THEREFORE; THESE *
* ESTIMATES MAY NOT REFLECT THE ACTUAL COSTS. *
* FOR PRICE INFORMATION SEE HELP COST *
**********

***NEW***
* ***PATENT NUMBERS (PN) AND BABS ACCESSION NUMBERS (BABSAN) CAN NOW BE SEARCHED, SELECTED AND TRANSFERRED.***
* ***NEW DISPLAY FORMATS ALLREF, ALLP AND BABSAN SHOW ALL REFERENCES, ALL PATENT REFERENCES, OR ALL BABS ACCESSION NUMBERS FOR A COMPOUND AT A GLANCE.***

FILE MARPAT
FILE CONTENT: 1961-PRESENT VOL 146 ISS 11 (20070309/ED)

SOME MARPAT RECORDS ARE DERIVED FROM INPI DATA FOR 1961-1987

MOST RECENT CITATIONS FOR PATENTS FROM MAJOR ISSUING AGENCIES
(COVERAGE TO THESE DATES IS NOT COMPLETE):

US 2007020715 25 JAN 2007
DE 102005032918 18 JAN 2007
EP 1743897 17 JAN 2007
JP 2007016265 25 JAN 2007
WO 2007012422 01 FEB 2007
GB 2427406 27 DEC 2006

FR 2888248 12 JAN 2007
RU 2291880 20 JAN 2007
CA 2551930 08 JAN 2007

Expanded G-group definition display now available.

FILE MEDLINE
FILE LAST UPDATED: 14 Mar 2007 (20070314/UP). FILE COVERS 1950 TO DATE.

All regular MEDLINE updates from November 15 to December 16 have been added to MEDLINE, along with 2007 Medical Subject Headings (MeSH(R)) and 2007 tree numbers.

The annual reload will be available in early 2007.

This file contains CAS Registry Numbers for easy and accurate substance identification.

FILE EMBASE
FILE COVERS 1974 TO 14 Mar 2007 (20070314/ED)

EMBASE is now updated daily. SDI frequency remains weekly (default) and biweekly.

This file contains CAS Registry Numbers for easy and accurate substance identification.

FILE BIOSIS
FILE COVERS 1969 TO DATE.
CAS REGISTRY NUMBERS AND CHEMICAL NAMES (CNs) PRESENT
FROM JANUARY 1969 TO DATE.

RECORDS LAST ADDED: 14 March 2007 (20070314/ED)

FILE DRUGU
FILE LAST UPDATED: 15 MAR 2007 <20070315/UP>
>>> DERWENT DRUG FILE (SUBSCRIBER) <<<

>>> FILE COVERS 1983 TO DATE <<<
>>> THESAURUS AVAILABLE IN /CT <<<

FILE WPIX
FILE LAST UPDATED: 14 MAR 2007 <20070314/UP>
MOST RECENT THOMSON SCIENTIFIC UPDATE: 200718 <200718/DW>
DERWENT WORLD PATENTS INDEX SUBSCRIBER FILE, COVERS 1963 TO DATE

>>> New reloaded DWPI Learn File (LWPI) available as well <<<

>>> YOU ARE IN THE NEW AND ENHANCED DERWENT WORLD PATENTS INDEX <<<

>>> New display format FRAGHITSTR available <<<
SEE ONLINE NEWS and
http://www.stn-international.de/archive/stn_online_news/fraghitstr_ex.pdf

>>> IPC Reform reclassification data for the backfile is being loaded into the database during January 2007.
There will not be any update date (UP) written for the reclassified documents, but they can be identified by 20060101/UPIC. <<<

FOR A COPY OF THE DERWENT WORLD PATENTS INDEX STN USER GUIDE,
PLEASE VISIT:
http://www.stn-international.de/training_center/patents/stn_guide.pdf

FOR DETAILS OF THE PATENTS COVERED IN CURRENT UPDATES, SEE
http://scientific.thomson.com/support/patents/coverage/latestupdates/

PLEASE BE AWARE OF THE NEW IPC REFORM IN 2006, SEE
http://www.stn-international.de/stndatabases/details/ipc_reform.html and
http://scientific.thomson.com/media/scpdf/ipcrdwpi.pdf

>>> FOR DETAILS ON THE NEW AND ENHANCED DERWENT WORLD PATENTS INDEX
PLEASE SEE
http://www.stn-international.de/stndatabases/details/dwpi_r.html <<<

FILE STNGUIDE
FILE CONTAINS CURRENT INFORMATION.
LAST RELOADED: Mar 9, 2007 (20070309/UP).

FILE CAOLD
FILE COVERS 1907-1966
FILE LAST UPDATED: 01 May 1997 (19970501/UP)

This file contains CAS Registry Numbers for easy and accurate substance identification. Title keywords, authors, patent assignees, and patent information, e.g., patent numbers, are now searchable from 1907-1966. TIFF images of CA abstracts printed between 1907-1966 are available in the PAGE display formats.

New CAS Information Use Policies, enter HELP USAGETERMS for details.

This file supports REG1stRY for direct browsing and searching of all substance data from the REGISTRY file. Enter HELP FIRST for more information.